Connecting via Winsock to STN

```
Welcome to STN International! Enter x:x
```

LOGINID: SSPTAJRK1626

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

```
Welcome to STN International
                 Web Page URLs for STN Seminar Schedule - N. America
NEWS
NEWS
                 "Ask CAS" for self-help around the clock
NEWS
                 New STN AnaVist pricing effective March 1, 2006
         FEB 27
NEWS 4
                 CA/CAplus enhanced with 1900-1906 U.S. patent records
         MAY 10
NEWS
         MAY 11
                KOREAPAT updates resume
NEWS
         MAY 19
                 Derwent World Patents Index to be reloaded and enhanced
NEWS
         MAY 30
                 IPC 8 Rolled-up Core codes added to CA/CAplus and
                 USPATFULL/USPAT2
NEWS
         MAY 30
                 The F-Term thesaurus is now available in CA/CAplus
      8
NEWS
         JUN 02
                 The first reclassification of IPC codes now complete in
                 INPADOC
NEWS 10
         JUN 26
                 TULSA/TULSA2 reloaded and enhanced with new search and
                 and display fields
NEWS 11
        JUN 28
                 Price changes in full-text patent databases EPFULL and PCTFULL
NEWS 12
        JUl 11
                 CHEMSAFE reloaded and enhanced
NEWS 13
        JUl 14 FSTA enhanced with Japanese patents
NEWS 14
        JUl 19 Coverage of Research Disclosure reinstated in DWPI
NEWS 15
       AUG 09 INSPEC enhanced with 1898-1968 archive
NEWS 16 AUG 28 ADISCTI Reloaded and Enhanced
        AUG 30
                CA(SM)/CAplus(SM) Austrian patent law changes
NEWS 17
NEWS 18 SEP 11
                CA/CAplus enhanced with more pre-1907 records
NEWS EXPRESS
             JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT
              MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
              AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.
NEWS HOURS
              STN Operating Hours Plus Help Desk Availability
NEWS LOGIN
              Welcome Banner and News Items
NEWS IPC8
              For general information regarding STN implementation of IPC 8
NEWS X25
              X.25 communication option no longer available
```

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

FILE 'HOME' ENTERED AT 15:38:13 ON 18 SEP 2006

=> file reg
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 15:38:24 ON 18 SEP 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 17 SEP 2006 HIGHEST RN 907180-17-0 DICTIONARY FILE UPDATES: 17 SEP 2006 HIGHEST RN 907180-17-0

New CAS Information Use Policies, enter HELP USAGETERMS for details.

Effective September 24, 2006, Concord 3D coordinates will no longer be available. Please contact CAS Customer Care (http://www.cas.org/supp.html) if you have a need for 3D coordinates.

TSCA INFORMATION NOW CURRENT THROUGH June 30, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

=>

Uploading C:\Program Files\Stnexp\Queries\10790288\Struc 1.str

Page 3

chain nodes : 7 8 15 16 17 18 19 20 21 22 23 24 27 28 ring nodes : 1 2 3 4 5 6 9 10 11 12 13 14 chain bonds : 6-7 7-8 7-15 8-9 8-17 15-16 16-27 17-18 17-24 18-19 19-20 20-21 21-22 22-23 27-28 ring bonds : 1-2 1-6 2-3 3-4 4-5 5-6 9-10 9-14 10-11 11-12 12-13 13-14 exact/norm bonds : 16-27 17-18 17-24 22-23 27-28 exact bonds : 6-7 7-8 7-15 8-9 8-17 15-16 18-19 19-20 20-21 21-22 normalized bonds : 1-2 1-6 2-3 3-4 4-5 5-6 9-10 9-14 10-11 11-12 12-13 13-14 isolated ring systems : containing 1 : 9 :

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:Atom 10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:CLASS 16:CLASS 17:CLASS 18:CLASS 19:CLASS 20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:CLASS 27:CLASS 28:CLASS

L1 STRUCTURE UPLOADED

=> d L1 HAS NO ANSWERS L1 STR

Structure attributes must be viewed using STN Express query preparation.

=> 11

SAMPLE SEARCH INITIATED 15:38:42 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED -1 TO ITERATE

100.0% PROCESSED 1 ITERATIONS 0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 1 TO 80 PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> 11 full

FULL SEARCH INITIATED 15:38:45 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED -45 TO ITERATE

100.0% PROCESSED 45 ITERATIONS 6 ANSWERS

SEARCH TIME: 00.00.01

6 SEA SSS FUL L1

=> file medline caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL

ENTRY SESSION FULL ESTIMATED COST 166.94 167.15

FILE 'MEDLINE' ENTERED AT 15:38:53 ON 18 SEP 2006

FILE 'CAPLUS' ENTERED AT 15:38:53 ON 18 SEP 2006

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

=> 13

L4 4 L3

=> d ibib abs hitstr 1-4

ANSWER 1 OF 4 CAPLUS COPYRIGHT 2006 ACS on STN

2006:14820 CAPLUS ACCESSION NUMBER:

144:260959

DOCUMENT NUMBER:

TITLE: Identification of a trace colored impurity in drug

substance by preparative liquid chromatography and

mass spectrometry

AUTHOR (S): Wang, Peng; Shi, Y.-J.; Helmy, Roy; Reamer, Robert;

Vailaya, Anant

Analytical Research, Merck Research Laboratories, CORPORATE SOURCE:

Rahway, NJ, 07065, USA

Rapid Communications in Mass Spectrometry (2005), SOURCE:

19(24), 3749-3754

CODEN: RCMSEF; ISSN: 0951-4198

John Wiley & Sons Ltd. PUBLISHER:

DOCUMENT TYPE: Journal

10790288.trn

English LANGUAGE:

6-(Nitrooxy)hexyl-(2z)-4-(acetyloxy)-3-[4-(methylsulfonyl)phenyl]-2phenylbut-2-enoate (enoate 1) was investigated as a novel therapy for pain relief. In a recent manufacturing run at the pilot plant scale, the enoate

drug

substance was found to have a yellowish color not observed previously. An unknown impurity at trace level was detected by high-performance liquid chromatog. (HPLC) anal. and found to be the primary cause for the color of the drug substance. The colored impurity was enriched by preparative HPLC and structurally elucidated by liquid chromatog./tandem mass spectrometry (LC/MS/MS). It was found that the colored impurity was derived from the product of oxidative dimerization of rofecoxib, an impurity present in the enoic acid intermediate. It was further revealed by the photodiode array and LC/MS/MS data that the colored impurity exists in the drug substance as a pair of double-bond isomers with one isomer at majority. These findings were also confirmed by synthesizing the colored impurity through the proposed pathway.

754241-98-0 IT

> RL: ANT (Analyte); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(identification of colored impurity in drug substance by preparative HPLC)

754241-98-0 CAPLUS RN

Benzeneacetic acid, α -[2-(acetyloxy)-1-[4-CN $(methylsulfonyl)phenyl]ethylidene]-, 6-(nitrooxy)hexyl ester, (<math>\alpha Z$)-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 20 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 2 OF 4 CAPLUS COPYRIGHT 2006 ACS on STN

2005:1315893 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 144:212486

Synthesis of a NO-Releasing Prodrug of Rofecoxib TITLE: Engelhardt, F. Conrad; Shi, Yao-Jun; Cowden, Cameron AUTHOR (S): J.; Conlon, David A.; Pipik, Brenda; Zhou, George;

McNamara, James M.; Dolling, Ulf-H. Department of Process Research, Merck Company, Rahway, CORPORATE SOURCE:

NJ, 07065-0900, USA

Journal of Organic Chemistry (2006), 71(2), 480-491 SOURCE:

CODEN: JOCEAH; ISSN: 0022-3263

American Chemical Society PUBLISHER:

DOCUMENT TYPE: Journal LANGUAGE: English

GT

AB A newly developed synthesis of a NO-releasing prodrug of rofecoxib is described. The highly productive process consists of five chemical steps and produces prodrug I in an overall 64% yield from com. available 3-phenyl-2-propyn-1-ol (II). The synthesis is highlighted by the carbometalation reaction of propargyl alc. II to generate the tetrasubstituted olefin core, sulfone acid III. Addnl., two alternate end-game strategies to prepare NO-COXIB I from this intermediate were explored and developed: (1) a convergent synthesis where a bromonitrate side chain is introduced in one step and (2) a two-step sequence that first installs the requisite six-carbon ester side chain followed by chemoselective nitration.

IT 754241-98-0P

MeO₂S

RL: SPN (Synthetic preparation); PREP (Preparation) (synthesis of a NO-releasing prodrug of rofecoxib in five chemical steps from 3-phenyl-2-propyn-1-ol)

RN 754241-98-0 CAPLUS

CN Benzeneacetic acid, α -[2-(acetyloxy)-1-[4-(methylsulfonyl)phenyl]ethylidene]-, 6-(nitrooxy)hexyl ester, (α Z)-(9CI) (CA INDEX NAME)

III

Double bond geometry as shown.

$$\begin{array}{c} \text{AcO} \\ \text{Z} \\ \text{O} \\ \text{O} \end{array} \begin{array}{c} \text{(CH2) 6} \\ \text{O} \end{array} \begin{array}{c} \text{NO2} \\ \text{O} \end{array}$$

REFERENCE COUNT: 14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2006 ACS on STN ACCESSION NUMBER: 2005:963804 CAPLUS

Page 7

DOCUMENT NUMBER: 143:266677

Process for making nitric oxide releasing prodrugs of TITLE: diaryl-2-(5H)-furanones as cyclooxygenase-2 inhibitors

Shi, Yao-Jun; Engelhardt, F. Conrad; Cowden, Cameron INVENTOR(S):

John; Conlon, David A.; Pipik, Brenda

PATENT ASSIGNEE(S): USA

U.S. Pat. Appl. Publ., 16 pp. SOURCE:

CODEN: USXXCO

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

DATE PATENT NO. KIND DATE APPLICATION NO. _____ --------------_____ 20050225 US 2005192346 **A1** 20050901 US 2005-66676 P 20040301 PRIORITY APPLN. INFO.: US 2004-549126P

CASREACT 143:266677; MARPAT 143:266677 OTHER SOURCE(S):

GI

The invention encompasses a novel process for making prodrugs of AB cyclooxygenase-2 selective inhibitors that convert in vivo to diaryl-2-(5H)-furanones and also liberate nitric oxide in vivo. As such, the compds. may be co-dosed with low-dose aspirin to treat chronic cyclooxygenase-2 mediated diseases or conditions, effectively reduce the risk of thrombotic cardiovascular events and potentially renal side effects and at the same time reduce the risk of GI ulceration or bleeding. I was prepared starting from 3-phenyl-2-propyn-1-ol and 4-thioanisole magnesium chloride, acetylation, and the intermediate converted to the carboxylic acid, the thio group oxidized to the methylsulfonyl derivative and reaction with 6-bromohexyl nitrate to give I.

Ι

IT 754241-98-0P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of nitric oxide releasing prodrugs of diaryl-2-(5H)-furanones as cyclooxygenase-2 inhibitors)

RN 754241-98-0 CAPLUS

Benzeneacetic acid, α -[2-(acetyloxy)-1-[4-CN

 $(methylsulfonyl)phenyl]ethylidene]-, 6-(nitrooxy)hexyl ester, (<math>\alpha Z$)-

(9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$AcO$$
 $CH_2)$ 6
 NO_2
 Ph

ANSWER 4 OF 4 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:739958 CAPLUS

DOCUMENT NUMBER: 141:260542

Preparation of nitric oxide releasing prodrugs of TITLE:

diaryl-2-(5H)-furanones as selective cyclooxygenase-2

inhibitors

Berthelette, Carl; Li, Lianhai; Sturino, Claudio; INVENTOR (S):

Wang, Zhaoyin

PATENT ASSIGNEE(S): Can.

SOURCE: U.S. Pat. Appl. Publ., 19 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.				
US 2004176331	A1 20040909	US 2004-790288				
AU 2004240700	A1 20041202	AU 2004-240700	20040301			
CA 2517490	AA 20041202	CA 2004-2517490	20040301			
WO 2004103955	A1 20041202	WO 2004-CA314	20040301			
W: AE, AG, AL,	AM, AT, AU, AZ,	BA, BB, BG, BR, BW,	BY, BZ, CA, CH,			
CN, CO, CR,	CU, CZ, DE, DK,	DM, DZ, EC, EE, EG,	ES, FI, GB, GD,			
		IN, IS, JP, KE, KG,				
		MD, MG, MK, MN, MW,				
		RO, RU, SC, SD, SE,				
		UG, US, UZ, VC, VN,				
		SD, SL, SZ, TZ, UG,	· · · · · · · · · · · · · · · · · · ·			
· · · · · · · · · · · · · · · · · · ·		AT, BE, BG, CH, CY,				
, , ,		IT, LU, MC, NL, PL,				
• • •	BJ, CF, CG, CI,	CM, GA, GN, GQ, GW,	ML, MR, NE, SN,			
. TD, TG						
EP 1601644	A1 20051207	EP 2004-761562	20040301			
R: AT, BE, CH,	DE, DK, ES, FR,	GB, GR, IT, LI, LU,	NL, SE, MC, PT,			
IE, SI, LT,	LV, FI, RO, MK,	CY, AL, TR, BG, CZ,	EE, HU, PL, SK			
PRIORITY APPLN. INFO.:		US 2003-452124P	P 20030305			
		WO 2004-CA314				
OTHER SOURCE(S):	MARPAT 141:2605					

GI

$$R^{1}$$
 R^{2}
 R^{3}
 R^{2}
 R^{3}
 R^{2}
 R^{3}
 R^{2}
 R^{3}
 R^{4}
 R^{2}
 R^{2}
 R^{3}
 R^{2}
 R^{3}

Me-SO₂ O-COCH₃ O-COCH₃ Z

II

Ι

AB Title compds. I [X = (CH2)n; n = 3-6; R1 = SO2Me, SO2NH2, SO2NHCOCF3, etc.; R2, R3 = H, halo, alkoxy, etc.; R4 = CO-alkyl, CO(CH2)mNR5R6; m = 1-4; R5, R6 = H, halo-substituted alkyl] and their pharmaceutically acceptable salts were prepared For example, O-alkylation of AgNO3 by bromide II (Z = Br), e.g., prepared from Rofecoxib in 6-steps, afforded nitrooxyhexyl II (Z = -ONO2). In human blood PGE2 inhibition production assays, nitrooxyhexyl II (Z = -ONO2) exhibited an IC50 value of 0.22 μM. Of note, the "unconverted prodrugs" of compds. I are inactive inhibitors of COX-1 and COX-2 activity. Compds. I are claimed useful for the treatment of cyclooxygenase-2 mediated diseases or conditions.

T 754241-98-0P 754241-99-1P 754242-00-7P

754242-01-8P 754242-02-9P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of nitric oxide releasing prodrugs of diarylfuranones as selective COX-2 inhibitors)

RN 754241-98-0 CAPLUS

CN Benzeneacetic acid, α -[2-(acetyloxy)-1-[4-(methylsulfonyl)phenyl]ethylidene]-, 6-(nitrooxy)hexyl ester, (α Z)-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 754241-99-1 CAPLUS

CN Glycine, (2Z)-2-[4-(methylsulfonyl)phenyl]-4-[[6-(nitrooxy)hexyl]oxy]-4-oxo-3-phenyl-2-butenyl ester, monohydrochloride (9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$NH_2$$
 Z
 $CCH_2)$ 6

 NO_2

HCl

RN 754242-00-7 CAPLUS

CN Benzeneacetic acid, α -[2-(acetyloxy)-1-[4-(methylsulfonyl)phenyl]ethylidene]-, 5-(nitrooxy)pentyl ester, (α Z)-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 754242-01-8 CAPLUS

CN Benzeneacetic acid, α -[2-(acetyloxy)-1-[4-(methylsulfonyl)phenyl]ethylidene]-, 7-(nitrooxy)heptyl ester, (α Z)-

Page 11

(9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$AcO$$
 $CCH_2)$ 7
 NO_2
 NO_2

RN 754242-02-9 CAPLUS

CN Glycine, (2Z)-2-[4-(methylsulfonyl)phenyl]-4-[[6-(nitrooxy)hexyl]oxy]-4-oxo-3-phenyl-2-butenyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

IT 754242-09-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of nitric oxide releasing prodrugs of diarylfuranones as selective COX-2 inhibitors)

RN 754242-09-6 CAPLUS

CN Glycine, N-[(1,1-dimethylethoxy)carbonyl]-, (2Z)-2-[4-(methylsulfonyl)phenyl]-4-[[6-(nitrooxy)hexyl]oxy]-4-oxo-3-phenyl-2-butenyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

=> file reg COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 21.41 188.56 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -3.00 -3.00

FILE 'REGISTRY' ENTERED AT 15:39:56 ON 18 SEP 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 17 SEP 2006 HIGHEST RN 907180-17-0 DICTIONARY FILE UPDATES: 17 SEP 2006 HIGHEST RN 907180-17-0

New CAS Information Use Policies, enter HELP USAGETERMS for details.

Effective September 24, 2006, Concord 3D coordinates will no longer be available. Please contact CAS Customer Care (http://www.cas.org/supp.html) if you have a need for 3D coordinates.

TSCA INFORMATION NOW CURRENT THROUGH June 30, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

=> rofecoxib

L5 1 ROFECOXIB

10790288.trn

=> d str

L5 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2006 ACS on STN

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

=> log h COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 6.58 195.14 SINCE FILE TOTAL DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SESSION ENTRY CA SUBSCRIBER PRICE 0.00 -3.00

SESSION WILL BE HELD FOR 60 MINUTES
STN INTERNATIONAL SESSION SUSPENDED AT 15:41:10 ON 18 SEP 2006

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:SSPTAJRK1626

PASSWORD:

* * * * * RECONNECTED TO STN INTERNATIONAL * * * * * * SESSION RESUMED IN FILE 'REGISTRY' AT 15:47:23 ON 18 SEP 2006 FILE 'REGISTRY' ENTERED AT 15:47:23 ON 18 SEP 2006 COPYRIGHT (C) 2006 American Chemical Society (ACS)

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
SINCE FILE TOTAL
ENTRY SESSION

Page 14

0.00 -3.00 CA SUBSCRIBER PRICE

=> d his

(FILE 'HOME' ENTERED AT 15:38:13 ON 18 SEP 2006)

FILE 'REGISTRY' ENTERED AT 15:38:24 ON 18 SEP 2006

L1 STRUCTURE UPLOADED

L20 L1

L3 6 L1 FULL

FILE 'MEDLINE, CAPLUS' ENTERED AT 15:38:53 ON 18 SEP 2006

L4

FILE 'REGISTRY' ENTERED AT 15:39:56 ON 18 SEP 2006

L5 1 ROFECOXIB

=> file medline caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL . ENTRY SESSION

FULL ESTIMATED COST 6.58 195.14

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL SESSION ENTRY

CA SUBSCRIBER PRICE 0.00 -3.00

FILE 'MEDLINE' ENTERED AT 15:47:38 ON 18 SEP 2006

FILE 'CAPLUS' ENTERED AT 15:47:38 ON 18 SEP 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

=> 15

L6 1523 L5

=> dup rem 16

PROCESSING COMPLETED FOR L6

1521 DUP REM L6 (2 DUPLICATES REMOVED)

=> 17 and prodrug

49 L7 AND PRODRUG

=> d ibib abs 1-49

ANSWER 1 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2006:740188 CAPLUS

DOCUMENT NUMBER:

145:159849

TITLE:

Methods and compositions using cyclooxygenase 2 (COX-2) inhibitors for the treatment of psychiatric

disorders, and combination therapies

INVENTOR(S):

Muller, Norbert

PATENT ASSIGNEE(S):

Germany

SOURCE:

U.S. Pat. Appl. Publ., 34 pp., Cont.-in-part of U.S.

Ser. No. 157,969.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

10790288.trn

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE		
US 2006167074	A1	20060727	US 2005-320757	20051230		
US 2003130334	A1	20030710	US 2002-157969	20020531		
EP 1627639	A2	20060222	EP 2005-24864	20020531		
R: AT, BE, CH,	DE, DK	, ES, FR,	GB, GR, IT, LI, LU, NL,	SE, MC, PT,		
IE, SI, LT,	LV, FI	, RO, MK,	CY, AL, TR			
PRIORITY APPLN. INFO.:			DE 2001-10129328	A 20010619		
			US 2002-364904P	P 20020314		
			US 2002-157969	A2 20020531		
		•	DE 2001-10129320	A 20010619		
•			EP 2002-738138	A3 20020531		

AB A method for the prevention, treatment, or inhibition of a psychiatric disorder, in particular schizophrenia, is described which comprises administering a COX-2 inhibitor or prodrug thereof to a subject.

Moreover, a method for the prevention, treatment, or inhibition of a psychiatric disorder, in particular schizophrenia or depressive disorders, is disclosed comprising administering to a subject a COX-2 inhibitor or prodrug thereof in combination with a neuroleptic drug or an antidepressant. Compns. and kits that are suitable for the practice of the method are also described.

L8 ANSWER 2 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2006:453900 CAPLUS

DOCUMENT NUMBER: 145:116702

TITLE: Racemic and chiral sulfoxides as potential prodrugs of

the COX-2 inhibitors Vioxx and Arcoxia

AUTHOR(S): Caturla, Francisco; Amat, Merce; Reinoso, Raquel F.;

Cordoba, Monica; Warrellow, Graham

CORPORATE SOURCE: Department of Medicinal Chemistry, Almirall

Prodesfarma S.A., Research Center, Barcelona, 08024,

Spain

SOURCE: Bioorganic & Medicinal Chemistry Letters (2006),

16(12), 3209-3212

CODEN: BMCLE8; ISSN: 0960-894X

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

GI

AB The preparation of the sulfoxide analogs (I) and (II), and their enantiomeric pure forms is discussed as well as their potential to act as prodrugs to the potent and selective sulfone-containing COX-2 inhibitors rofecoxib and etoricoxib. Sulfoxides I and II were shown to be effectively transformed

Page 16

in vivo into rofecoxib and etoricoxib, resp., after oral administration in rats. In the case of sulfoxide I, both a slightly improved pharmacokinetic profile and a better pharmacol. activity in an arthritis model were seen when compared with rofecoxib.

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 3 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

6

ACCESSION NUMBER:

2005:1315893 CAPLUS

DOCUMENT NUMBER:

144:212486

TITLE:

Synthesis of a NO-Releasing Prodrug of

Rofecoxib

AUTHOR (S):

Engelhardt, F. Conrad; Shi, Yao-Jun; Cowden, Cameron J.; Conlon, David A.; Pipik, Brenda; Zhou, George;

McNamara, James M.; Dolling, Ulf-H.

CORPORATE SOURCE:

Department of Process Research, Merck Company, Rahway,

NJ, 07065-0900, USA

SOURCE:

Journal of Organic Chemistry (2006), 71(2), 480-491

CODEN: JOCEAH; ISSN: 0022-3263

PUBLISHER:

American Chemical Society

DOCUMENT TYPE:

Journal

LANGUAGE:

English

GI

14

AB A newly developed synthesis of a NO-releasing prodrug of rofecoxib is described. The highly productive process consists of five chemical steps and produces prodrug I in an overall 64% yield from com. available 3-phenyl-2-propyn-1-ol (II). The synthesis is highlighted by the carbometalation reaction of propargyl alc. II to generate the tetrasubstituted olefin core, sulfone acid III. Addnl., two alternate end-game strategies to prepare NO-COXIB I from this intermediate were explored and developed: (1) a convergent synthesis where a bromonitrate side chain is introduced in one step and (2) a two-step sequence that first installs the requisite six-carbon ester side chain followed by chemoselective nitration.

REFERENCE COUNT:

THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 4 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:1294044 CAPLUS

DOCUMENT NUMBER: 144:17160

TITLE: Method using camptothecin compounds, pyrimidine

derivatives, and antitumor agents for treating

abnormal cell growth

INVENTOR(S): Denis, Louis J.; Compton, Linda D.

PATENT ASSIGNEE(S): Pfizer Inc, USA

SOURCE: U.S. Pat. Appl. Publ., 32 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.			DATE			
	'					
US 2005272755	A1 20051208	US 2005-145097	20050603			
WO 2005117980	A1 20051215	WO 2005-IB1527	20050523			
W: AE, AG, AL,	AM, AT, AU, AZ,	BA, BB, BG, BR, BW, BY,	BZ, CA, CH,			
CN, CO, CR,	CU, CZ, DE, DK,	DM, DZ, EC, EE, EG, ES,	FI, GB, GD,			
GE, GH, GM,	HR, HU, ID, IL,	IN, IS, JP, KE, KG, KM,	KP, KR, KZ,			
LC, LK, LR,	LS, LT, LU, LV,	MA, MD, MG, MK, MN, MW,	MX, MZ, NA,			
NG, NI, NO,	NZ, OM, PG, PH,	PL, PT, RO, RU, SC, SD,	SE, SG, SK,			
SL, SM, SY,	TJ, TM, TN, TR,	TT, TZ, UA, UG, US, UZ,	VC, VN, YU,			
ZA, ZM, ZW						
RW: BW, GH, GM,	KE, LS, MW, MZ,	NA, SD, SL, SZ, TZ, UG,	ZM, ZW, AM,			
AZ, BY, KG,	KZ, MD, RU, TJ,	TM, AT, BE, BG, CH, CY,	CZ, DE, DK,			
EE, ES, FI,	FR, GB, GR, HU,	IE, IS, IT, LT, LU, MC,	NL, PL, PT,			
RO, SE, SI,	SK, TR, BF, BJ,	CF, CG, CI, CM, GA, GN,	GQ, GW, ML,			
MR, NE, SN,	TD, TG					

PRIORITY APPLN. INFO.:

US 2004-577268P

P 20040604

The invention discloses a method for treating abnormal cell growth in a subject, comprising administering to the subject (a) a compound selected from a camptothecin, a camptothecin derivative, or a pharmaceutically acceptable salt, solvate or prodrug thereof; (b) a pyrimidine derivative or a pharmaceutically acceptable salt, solvate or prodrug thereof; and (c) an antitumor agent selected from antiproliferative agents, kinase inhibitors, angiogenesis inhibitors, growth factor inhibitors, COX-1 inhibitors, COX-2 inhibitors, mitotic inhibitors, alkylating agents, antimetabolites, intercalating antibiotics, growth factor inhibitors, radiation, cell cycle inhibitors, enzymes, topoisomerase inhibitors, biol. response modifiers, antibodies, cytotoxics, antihormones, antiandrogens and combinations thereof.

```
L8 ANSWER 5 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN
```

ACCESSION NUMBER: 2005:1291841 CAPLUS

DOCUMENT NUMBER: 144:40800

TITLE: Glucosamine and glucosamine/anti-inflammatory mutual

prodrugs, compositions, and methods

INVENTOR(S): Capomacchia, Anthony C.; Garner, Solomon T., Jr.;

Beach, J. Warren

PATENT ASSIGNEE(S): The University of Georgia Research Center Inc., USA

SOURCE: PCT Int. Appl., 83 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

```
KIND
                               DATE
                                         APPLICATION NO.
     PATENT NO.
                                                                DATE
                        ____
                               ------
                                          -----
     -----
                                                                 -----
                                        WO 2005-US11739
                         A2
                               20051208
                                                                 20050407
     WO 2005116086
                        A3
                               20060824
     WO 2005116086
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
            LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
            NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL,
            SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA,
            ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
            AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
            EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT,
            RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
            MR, NE, SN, TD, TG
                                           US 2004-560128P
PRIORITY APPLN. INFO.:
                                                              P 20040407
                        MARPAT 144:40800
OTHER SOURCE(S):
    Mutual prodrugs of glucosamine, and derivs. and analogs of glucosamine and
     an anti-inflammatory agent, compns. thereof, and methods for, e.g.,
     treating disorders and conditions by administration of the compns. are
    provided. Topical compns. of glucosamine, and derivs. and analogs of
    glucosamine are also provided.
    ANSWER 6 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:
                        2005:1155282 CAPLUS
DOCUMENT NUMBER:
                        143:427372
TITLE:
                        Methods and compositions for preventing or treating
                        periodontal diseases using, for example, Resolvin El
INVENTOR(S):
                        Van, Dyke Thomas E.; Petasis, Nicos A.; Serhan,
                        Charles N.
PATENT ASSIGNEE(S):
                        USA
SOURCE:
                        U.S. Pat. Appl. Publ., 19 pp.
                        CODEN: USXXCO
DOCUMENT TYPE:
                        Patent
                        English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
    PATENT NO.
                        KIND
                               DATE
                                         APPLICATION NO.
                                                                 DATE
     -----
                        ____
                               -----
                                          -----
                                                                 _____
                        A1
                               20051027
                                         US 2005-106141
                                                                 20050414
    US 2005238589
                        A1
                               20051110
                                          WO 2005-US12552
                                                                 20050414
    WO 2005105025
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
            LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
            NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL,
            SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA,
            ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
            AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
            EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT,
            RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
            MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                           US 2004-562099P P 20040414
```

MARPAT 143:427372

GΙ

OTHER SOURCE(S):

$$\mathbb{R}^2$$
 \mathbb{R}^3
 \mathbb{R}^4
 \mathbb{R}^4
 \mathbb{R}^1

AB Methods and compns. for preventing or treating periodontal diseases, including gingivitis and periodontitis are provided. The compns. comprise a prophylactically or therapeutically effective amount of a compound I (R1,R2, R3 = OR, OX1, SR, SX2, N(R)2, NHX3, NRC(O)R, NRC(O)N(R)2, CO2R, C(O)N(R)2,SO2R, NRSO2, C(O)R, SO2N(R)2; R = C1-6 aliphatic, 3-8 membered saturated, aryl; heterocycle, heteroaryl; X1,X2,X3 = protecting group; R4 = NRC(0)R, NRC(O)N(R), C(O)OR, C(O)N(R)2, SO2R, NRSO2R, C(O)R, or SO2N(R)2), or a pharmaceutically acceptable salt or prodrug thereof and a pharmaceutically acceptable carrier. The composition further includes a COX-2 inhibitor selected from celecoxib, rofecoxib, and valdecoxib. The invention also provides methods for preventing or treating secondary diseases within or beyond the oral cavity that are related to periodontal disease, such as cardiovascular diseases, pregnancy complications, and diabetes. Thus, topical delivery of Resolvin El suspended in ethanol (7 μg/mL) every other day for 6 wk prevented both the bone loss and inflammatory changes in rabbits treated either with ligature alone or ligature plus topical Porphyromonas gingivalis (model of periodontal disease).

Ι

ANSWER 7 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN L8

2005:824492 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 143:222525

Method of using 3-cyano-4-arylpyridine derivatives as TITLE:

modulators of androgen receptor function, preparation

thereof, and use with other agents

INVENTOR(S): Nirschl, Alexandra A.; Hamann, Lawrence G.

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 25 pp.

Patent

CODEN: USXXCO

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

DOCUMENT TYPE:

PATENT NO. KIND DATE APPLICATION NO. DATE ______ ______ ----_____ _____ US 2005182105 US 2005-48437 20050201 **A1** 20050818 PRIORITY APPLN. INFO.: US 2004-541780P P 20040204

OTHER SOURCE(S): MARPAT 143:222525

GT

$$\begin{array}{c|c}
G \\
R1 \\
R2X \\
NR3R4 \\
I
\end{array}$$

AB A method is provided for treating androgen receptor-associated conditions, such as age-related diseases, e.g. sarcopenia, employing a compound I [R1 = CN, H; X = O, S; R2 = (substituted) alkyl, (substituted) cycloalkyl, etc; R3, R4 = H, (substituted) alkyl, etc.; G = (substituted) aryl, (substituted) heteroaryl], or a pharmaceutically acceptable salt or prodrug ester thereof. Preparation of selected I is described. I may be used in combination with other agents.

L8 ANSWER 8 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:696865 CAPLUS

DOCUMENT NUMBER: 143:193802

TITLE: Preparation of nitric oxide releasing prodrugs of

diaryl-2(5H)-furanones as cyclooxygenase-2 inhibitors

INVENTOR(S): Berthelette, Carl; Li, Lianhai; Beaulieu, Christian;

Wang, Zhaoyin; Sturino, Claudio F.

PATENT ASSIGNEE(S): Merck Frosst Canada & Co., Can.

SOURCE: PCT Int. Appl., 41 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICAT	TION NO. DATE
WO 2005070874 A1 20050804 WO 2005-	-CA84 20050125
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG,	, BR, BW, BY, BZ, CA, CH,
CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC,	, EE, EG, ES, FI, GB, GD,
GE, GH, GM, HR, HU, ID, IL, IN, IS, JP,	, KE, KG, KP, KR, KZ, LC,
LK, LR, LS, LT, LU, LV, MA, MD, MG, MK,	, MN, MW, MX, MZ, NA, NI,
NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC,	, SD, SE, SG, SK, SL, SY,
TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ,	, VC, VN, YU, ZA, ZM, ZW
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL,	, SZ, TZ, UG, ZM, ZW, AM;
AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE,	, BG, CH, CY, CZ, DE, DK,
EE, ES, FI, FR, GB, GR, HU, IE, IS, IT,	, LT, LU, MC, NL, PL, PT,
RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI,	, CM, GA, GN, GQ, GW, ML,
MR, NE, SN, TD, TG	
PRIORITY APPLN. INFO.: US 2004	-540101P P 20040127

OTHER SOURCE(S): MARPAT 143:193802

GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Title compds. I [n = 1-6; R1 = SO2CH3, SO2NH2; R2-3 = H, halo, alkoxy, etc.; R4 = alkyl, Ph, etc.] are prepared For instance, II is prepared in several steps from 4-(4-(methanesulfonyl)phenyl)-3-phenyl-5H-furan-2-one and hex-5-en-1-ol. I are nitric oxide-releasing prodrugs of

Page 21

diary1-2(5H)-furanones useful in the treatment of cyclooxygenase-2 mediated diseases [no data]. I may also be used as a combination therapy with low-dose aspirin to treat chronic cyclooxygenase-2 mediated diseases or conditions while also reducing the risk of thrombotic cardiovascular events.

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 9 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:460617 CAPLUS

DOCUMENT NUMBER: 144:186912

TITLE: Examination of 209 drugs for inhibition of cytochrome

P450 2C8

AUTHOR(S): Walsky, Robert L.; Gaman, Emily A.; Obach, R. Scott

CORPORATE SOURCE: Pharmacokinetics, Pharmacodynamics, and Drug

Metabolism, Pfizer Global Research and Development, Groton/New London Laboratories, Groton, CT, USA

SOURCE: Journal of Clinical Pharmacology (2005), 45(1), 68-78

CODEN: JCPCBR: ISSN: 0091-2700

PUBLISHER: Sage Publications

DOCUMENT TYPE: Journal LANGUAGE: English

Cytochrome P 450 2C8 is involved in the metabolism of drugs such as paclitaxel, repaglinide, rosiglitazone, and cerivastatin, among others. An in vitro assessment of 209 frequently prescribed drugs and related xenobiotics was carried out to examine their potential to inhibit CYP2C8. A validated sensitive, moderate-throughput high-performance liquid chromatog./tandem mass spectrometry(HPLC/MS/MS) assay was used to detect N-desethylamodiaguine, the CYP2C8-derived major metabolite of amodiaguine metabolism, using heterologously expressed recombinant CYP2C8 (rhCYP2C8) and pooled human liver microsomes. The 209 drugs were first tested at 30 uM for their ability to inhibit rhCYP2C8. Forty-eight compds. exhibited greater than 50% inhibition and were further evaluated for measurement of IC50. The six most potent inhibitors (IC50 <1 μ M) from this set were measured for IC50 in pooled human liver microsomes, and the most potent inhibitor identified was the leukotriene receptor antagonist, montelukast (IC50 = 19.6 nM). Inhibitors of CYP2C8 were identified from a wide variety of therapeutic classes, with no single class predominating. Other potent inhibitors included candesartan cilexetil (cyclohexylcarbonate ester prodrug of candesartan), zafirlukast, clotrimazole, felodipine, and mometasone furoate. Seventeen moderate inhibitors of rhCYP2C8 (1 < IC50 < 10 μM) included salmeterol, raloxifene, fenofibrate, ritonavir, levothyroxine, tamoxifen, loratadine, quercetin, oxybutynin, medroxyprogesterone, simvastatin, ketoconazole, ethinyl estradiol, spironolactone, lovastatin, nifedipine, and irbesartan. These in vitro data were used along with clin. pharmacokinetic information in predicting potential drug-drug interactions that could occur by inhibition of CYP2C8. Although almost all drugs tested are not expected to cause drug interactions via inhibition of CYP2C8, montelukast was identified as being of concern as a potential inhibitor of clin. These findings are discussed in context to potential drug interactions that could be observed between these agents and drugs for which CYP2C8 is involved in metabolism and warrant investigation of the possibility of clin. drug interactions mediated by inhibition of this enzyme.

REFERENCE COUNT: 38 THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 10 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:451140 CAPLUS

DOCUMENT NUMBER: 142:476264

TITLE:

Compositions of a cyclooxygenase-2 selective inhibitor and a neurotrophic factor-modulating agent for the treatment of central nervous system-mediated disorders

INVENTOR(S):

Taylor, Duncan P.; Stephenson, Diane T. Pharmacia & Upjohn Company LLC, USA

PATENT ASSIGNEE(S):

SOURCE:

PCT Int. Appl., 153 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA	PATENT NO.								APPLICATION NO.						DATE			
			15					WO 2004-US37882						20041112				
							AU,		BA.	BB.	BG	BR.	BW.	BY.	B7.	CA.	CH.	
	***	•	•	•	•		DE,	•	•	•	•	-	-	•	•	•	•	
		•	•	•	•		ID,	•	•	•	-		-		•	•	•	
		•	•	•	•	•	LV,	•	•	•			•	•	•	•	•	
		•		•		-	PL,		-	•	-	-	-		-		•	
		•	•		•	•	TZ,	•	•	•	•	•	•	•		•	•	
	DW.						MW,		-	-	-	-	_		-			
	RW:																	
							RU,											
		•	•	•	•	•	GR,	•		•		•	-	•	•			
		•	•	•	•	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	Gw,	МП,	MR,	
	05.45	•	SN,	•			0005	0506		a					-	0041		
	2545						2005			-								
	2005						2005											
EP	1684						2006											
	R:	•	•	-	•		ES,	•	-	•	•		-					
					LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	HU,	PL,	SK,	
		HR,	IS,	YU														
PRIORIT	Y APP	LN.	INFO	.:											P 2			
									1	WO 2	004-1	JS37	882	1	W 2	0041	112	

OTHER SOURCE(S): MARPAT 142:476264

The invention provides compns. and methods for the treatment of central nervous system-mediated disorders. More particularly, the invention provides a combination therapy for the treatment of a central nervous system-mediated disorder which comprises the administration of a neurotrophic factor-modulating agent in combination with a cyclooxygenase-2 selective inhibitor.

ANSWER 11 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2005:409223 CAPLUS

DOCUMENT NUMBER:

142:441891

TITLE:

Method and compositions for the treatment and prevention of pain and inflammation with

cyclooxygenase-2 inhibitors and polyunsaturated fatty

acids

INVENTOR (S):

Pulaski, Steven P.; Kundel, Susan

PATENT ASSIGNEE(S):

Pharmacia Corporation, USA

SOURCE:

U.S. Pat. Appl. Publ., 61 pp., Cont.-in-part of U.S.

Ser. No. 215,539.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2005101563	A1	20050512	US 2004-783160	20040219
US 2003114416	A1	20030619	US 2002-215539	20020809
CN 1575182	A	20050202	CN 2002-820121	20020813
ZA 2004001163	Α	20050622	ZA 2004-1163	20040212
PRIORITY APPLN. INFO.:			US 2001-312211P F	20010814
			US 2002-215539 A	2 20020809

AB A method of preventing or treating pain or inflammation in a subject is provided by administering to the subject a Cox-2 inhibitor and a polyunsatd. fatty acid, or a prodrug thereof, wherein the amount of a Cox-2 inhibitor and polyunsatd. fatty acid or a pharmaceutically acceptable salt or prodrug thereof together constitute a pain or inflammation suppressing treatment or prevention effective amount Glucosamine and/or chondroitin can optionally be present. Therapeutic compns. that contain the combination of Cox-2 inhibitor and polyunsatd. fatty acid and, optionally, the glucosamine and/or chondroitin, are disclosed, as are pharmaceutical compns.

L8 ANSWER 12 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2005:259661 CAPLUS

DOCUMENT NUMBER:

142:336520

TITLE:

Preparation, pharmaceutical compositions, and methods comprising combinations of 2-alkylidene-19-nor-vitamin

D derivatives and a cyclooxgenase-2 inhibitor

INVENTOR(S):

Thompson, David D.

PATENT ASSIGNEE(S): SOURCE:

Pfizer Inc., USA
U.S. Pat. Appl. Publ., 20 pp.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA	PATENT NO.				KIND DATE			APPLICATION NO.						DATE			
US	2005	0651	30		A1	_	2005	0324	1	US 2	004-	9435	61		21	0040	916
WO	2005	0279	18		A1 20050331			WO 2004-IB2913						20040906			
	W: -	ΑE,	AG,	AL,	AM,	ΑT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	ΚZ,	LC,
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,
							PL,										
		TJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW
	RW:	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,
		AZ,	BY,	KG,	KZ,	MD,	RU,	ТJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,
		EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,
		SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,
		SN,	TD,	TG													
PRIORITY	Y APP	LN.	INFO	. :					1	US 2	003-	5040	03P	1	P 20	0030	919

GI

The invention relates to pharmaceutical compns., and methods of treatment AB comprising administering to a patient in need of a combination of a 2-alkylidene-19-nor-vitamin D derivative and a cyclooxgenase-2 inhibitor, or a pharmaceutically acceptable salt or prodrug thereof. Particularly, the invention relates to pharmaceutical compns. and methods of treatment comprising administering to a patient in need of 2-methylene-19-nor-20(S)-10,25-dihydroxyvitamin D3 and a cyclooxgenase-2 inhibitor, or a pharmaceutically acceptable salt or prodrug thereof. Thus, 1a,25-dihydroxy-2-methylene-19-norvitamin D3 (I) was prepared in 11 steps from (-)-quinic acid. and (20S)-1α,25-dihydroxy-2methylene-19-norvitamine D3 was prepared from (20S)-25-[(triethylsilyl)oxy]des-A,B-cholestan-8-one in 4 steps.

ANSWER 13 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:216610 CAPLUS

DOCUMENT NUMBER: 142:291412

Compositions of a cyclooxygenase-2 selective inhibitor TITLE:

> and a corticotropin releasing factor antagonist for the treatment of ischemic-mediated central nervous

system disorders or injury

Arneric, Stephen P. INVENTOR(S):

PATENT ASSIGNEE(S): Pharmacia Corporation, USA SOURCE:

PCT Int. Appl., 155 pp.

Ι

CODEN: PIXXD2

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE
		·	
WO 2005020910	A2 200503	LO WO 2004-US27600	20040826
WO 2005020910 .	A3 2005060)9	
W: AE, AG, AL,	AM, AT, AU, A	Z, BA, BB, BG, BR, BW, E	BY, BZ, CA, CH,
CN, CO, CR,	CU, CZ, DE, DI	K, DM, DZ, EC, EE, EG, E	ES, FI, GB, GD,
GE, GH, GM,	HR, HU, ID, II	L, IN, IS, JP, KE, KG, K	CP, KR, KZ, LC,
LK, LR, LS,	LT, LU, LV, M	A, MD, MG, MK, MN, MW, M	IX, MZ, NA, NI,
NO, NZ, OM,	PG, PH, PL, PT	r, RO, RU, SC, SD, SE, S	SG, SK, SL, SY,
TJ, TM, TN,	TR, TT, TZ, UZ	A, UG, US, UZ, VC, VN, Y	TU, ZA, ZM, ZW

```
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
             SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
             SN, TD, TG
                                20050421
     US 2005085479
                          A1
                                            US 2004-926751
                                                                   20040826
                                            US 2003-498148P
                                                                P 20030827
PRIORITY APPLN. INFO.:
                         MARPAT 142:291412
OTHER SOURCE(S):
     The invention provides compns. and methods for the treatment of
     ischemic-mediated central nervous system disorder or injury. More
     particularly, the invention provides a combination therapy for the
     treatment of a central nervous system ischemic-mediated disorder or injury
     comprising the administration to a subject of a cyclooxygenase-2 selective
     inhibitor and a corticotropin releasing factor antagonist or a
     pharmaceutically acceptable salt or a prodrug thereof.
     ANSWER 14 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:
                         2005:177827 CAPLUS
DOCUMENT NUMBER:
                         142:254634
                         Compositions of a cyclooxygenase-2 selective inhibitor
TITLE:
                         and a serotonin-modulating agent for the treatment of
                         central nervous system damage
INVENTOR(S):
                         Stephenson, Diane T.
PATENT ASSIGNEE(S):
                         Pharmacia Corporation, USA
SOURCE:
                         PCT Int. Appl., 172 pp.
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                       KIND
                               DATE
                                          APPLICATION NO.
                                                                  DATE
     ------
                         ----
                                -----
                                            -----
                                                                   -----
                                           WO 2004-US22059
     WO 2005018541
                         A2
                                20050303
                                                                   20040708
     WO 2005018541
                         A3
                                20060309
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
             SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
             SN, TD, TG
                                20050414
                                            US 2004-887112
     US 2005080084
                         A1
                                                                   20040708
                                            US 2003-486549P
PRIORITY APPLN. INFO.:
                                                                P 20030711
                        MARPAT 142:254634
OTHER SOURCE(S):
     The invention provides compns. and methods for the treatment of central
     nervous system damage in a subject. More particularly, the invention
```

AB The invention provides compns. and methods for the treatment of central nervous system damage in a subject. More particularly, the invention provides a combination therapy for the treatment of a central nervous system ischemic condition or a central nervous system traumatic injury comprising the administration to a subject of a serotonin-modulating agent in combination with a cyclooxygenase-2 selective inhibitor.

L8 ANSWER 15 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:99157 CAPLUS

DOCUMENT NUMBER: 142:170033

TITLE: Methods and compositions for the treatment or

prevention of human immunodeficiency virus and related conditions using cyclooxygenase-2 selective inhibitors

and antiviral agents

INVENTOR(S):

Maziasz, Timothy

PATENT ASSIGNEE(S):

USA

SOURCE:

U.S. Pat. Appl. Publ., 172 pp.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE		
US 2005026902	A1	20050203	US 2004-769485	20040130		
PRIORITY APPLN. INFO.:			US 2003-443910P P	20030131		
OTHER SOURCE(S):	MARPAT	142:170033				

The present invention provides compns. and methods for the treatment of human immunodeficiency virus (HIV) infection as well as HIV associated diseases and related disorders. More particularly, the invention provides a combination therapy for the treatment of HIV infection as well as HIV associated diseases and related disorders comprising the administration to a subject of an anti-human immunodeficiency virus agent in combination with a cyclooxygenase-2 selective inhibitor or an isomer or a pharmaceutically acceptable salt, ester, or prodrug thereof.

ANSWER 16 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2005:76247 CAPLUS

DOCUMENT NUMBER:

142:148812

TITLE:

Compositions of a cyclooxygenase-2 selective inhibitor

and a non-NMDA glutamate modulator for the treatment

of central nervous system damage

INVENTOR(S):

Stephenson, Diane T.; Taylor, Duncan P.

PATENT ASSIGNEE(S): SOURCE:

Pharmacia Corporation, USA PCT Int. Appl., 150 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	PATENT NO.						DATE										
						-									-		- .
WO	2005	0071	06		A2		2005	20050127 WO 2004-US22189				20040708					
WO	2005	0071	06		A 3		2006	0608									
	W:	ΑE,	AG,	AL,	AM,	ΑT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KΕ,	KG,	ΚP,	KR,	ΚZ,	LC,
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	ΜZ,	NA,	NI,
		NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,
		ТJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UΖ,	VC,	VN,	YU,	ZA,	ZM,	ZW
	RW:	BW,	GH,	GM,	KΕ,	LS,	MW,	MZ,	NA,	SD,	ŞL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,
		ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM,	ΑT,	ВE,	BG,	CH,	CY,	CZ,	DE,	DK,
		EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,
		SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,
		SN,	TD,	TG													
ບຣ	2005	1015	97		A1		2005	0512	1	US 2	004-	8870	35		2	0040	708
PRIORITY	Y APP	LN.	INFO	. :					1	US 2	003-	4866	54P]	P 20	0030'	710
OTHER SO	OURCE	(S):			MAR	PAT	142:	1488	12								
AB The invention provides compns.								and 1	meth	ods	for	the '	trea	tment	t of	cent	tral

nervous system damage in a subject. More particularly, the invention provides a combination therapy for the treatment of a central nervous system ischemic condition or a central nervous system traumatic injury comprising the administration to a subject of a non-NMDA glutamate modulator in combination with a cyclooxygenase-2 selective inhibitor.

L8 ANSWER 17 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:1020415 CAPLUS

DOCUMENT NUMBER: 142:190038

TITLE: Selective cyclooxygenase-2 inhibitors: similarities

and differences

AUTHOR(S): Brune, K.; Hinz, B.

CORPORATE SOURCE: Department of Experimental and Clinical Pharmacology

and Toxicology, Emil Fischer Center, Friedrich

Alexander University, Erlangen, Germany

SOURCE: Scandinavian Journal of Rheumatology (2004), 33(1),

1-6

CODEN: SJRHAT; ISSN: 0300-9742

PUBLISHER: Taylor & Francis

DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

A review. The enzyme cyclooxygenase (COX) was shown to exist as two AB distinct isoforms about a decade ago. COX-1 is constitutively expressed as a 'housekeeping' enzyme in nearly all tissues, and mediates physiol. responses (e.g. cytoprotection of the stomach, and platelet aggregation). On the other hand, COX-2, expressed by cells involved in inflammation (e.g. macrophages, monocytes, synoviocytes), has emerged as the isoform that is primarily responsible for the synthesis of prostanoids involved in acute and chronic inflammatory states. Consequently, the hypothesis that selective inhibition of COX-2 might have therapeutic actions similar to those of non-steroidal anti-inflammatory drugs, but without causing gastrointestinal side effects, was the rationale for the development of selective inhibitors of the COX-2 isoenzyme. Selective COX-2 inhibitors currently used in the clinic are the sulfonamides celecoxib and valdecoxib (parecoxib is a prodrug of valdecoxib), as well as the methylsulfones rofecoxib and etoricoxib. Furthermore, the phenylacetic acid derivative lumiracoxib has gained permission recently to be marketed in Europe. This review discusses the clin. relevant similarities and differences of these substances, with particular emphasis on their diverse pharmacokinetic characteristics.

REFERENCE COUNT: 47 THERE ARE 47 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 18 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2004:754407 CAPLUS

DOCUMENT NUMBER:

141:271579

TITLE:

Treatment and prevention of obesity with COX-2 inhibitors alone or in combination with weight-loss

agents

INVENTOR(S):

Briggs, Michael; Ornberg, Richard; Hauser, Scott;

Koki, Alane

PATENT ASSIGNEE(S):

Pharmacia Corporation, USA

PCT Int. Appl., 180 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.

KIND DATE

APPLICATION NO.

DATE

```
----
                              -----
                                          -----
     -----
    WO 2004078113
                        A2
                              20040916 WO 2004-US3219
                                                                20040205
                        A3 20051013
    WO 2004078113
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE,
            BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU,
            MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN,
            GQ, GW, ML, MR, NE, SN, TD, TG
                                          US 2004-773019
    US 2004204472
                        A1
                              20041014
                                                                20040205
                                          US 2004-773019 20040205
US 2003-451885P P 20030304
PRIORITY APPLN. INFO.:
    A method for preventing or treating obesity and obesity-related
    complications in a subject involves a monotherapy with a Cox-2 inhibitor
    or a combination therapy with a Cox-2 inhibitor and a conventional
    weight-loss agent. Also described are therapeutic compns. comprising a Cox-2
     inhibitor and a conventional weight-loss agent. Pharmaceutical compns. and
    kits for implementing the present method are also described.
    ANSWER 19 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:
                        2004:739958 CAPLUS
DOCUMENT NUMBER:
                        141:260542
TITLE:
                        Preparation of nitric oxide releasing prodrugs of
                        diary1-2-(5H)-furanones as selective cyclooxygenase-2
                        inhibitors
INVENTOR(S):
                        Berthelette, Carl; Li, Lianhai; Sturino, Claudio;
                        Wang, Zhaoyin
PATENT ASSIGNEE(S):
                        Can.
                        U.S. Pat. Appl. Publ., 19 pp.
SOURCE:
                        CODEN: USXXCO
DOCUMENT TYPE:
                        Patent
                        English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                     KIND DATE
    PATENT NO.
                                        APPLICATION NO.
                                                               DATE
                      ---- . ------
                                        -----
                                                                -----
                                        US 2004-790288 20040301
    US 2004176331
                       A1 20040909
    AU 2004240700
                       Al.
                              20041202
                                        AU 2004-240700
    CA 2517490
                       AA
                              20041202
                                         CA 2004-2517490
                                                               20040301
                                         WO 2004-CA314
    WO 2004103955
                       A1
                              20041202
                                                                20040301
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
            TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
            BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
            ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
            SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
            TD, TG
                                         EP 2004-761562
                                                                 20040301
    EP 1601644
                         A1
                              20051207
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK
                                          US 2003-452124P P 20030305
PRIORITY APPLN. INFO.:
```

W 20040301

WO 2004-CA314

OTHER SOURCE(S): MARPAT 141:260542

GΙ

$$R^{1}$$
 R^{2}
 R^{2}
 R^{3}
 R^{3}
 R^{2}
 R^{3}
 R^{4}
 R^{2}
 R^{2}
 R^{3}
 R^{4}
 R^{2}
 R^{2}
 R^{3}
 R^{4}
 R^{4

AB Title compds. I [X = (CH2)n; n = 3-6; R1 = SO2Me, SO2NH2, SO2NHCOCF3, etc.; R2, R3 = H, halo, alkoxy, etc.; R4 = CO-alkyl, CO(CH2)mNR5R6; m = 1-4; R5, R6 = H, halo-substituted alkyl] and their pharmaceutically acceptable salts were prepared For example, O-alkylation of AgNO3 by bromide II (Z = Br), e.g., prepared from Rofecoxib in 6-steps, afforded nitrooxyhexyl II (Z = -ONO2). In human blood PGE2 inhibition production assays, nitrooxyhexyl II (Z = -ONO2) exhibited an IC50 value of 0.22 μM. Of note, the "unconverted prodrugs" of compds. I are inactive inhibitors of COX-1 and COX-2 activity. Compds. I are claimed useful for the treatment of cyclooxygenase-2 mediated diseases or conditions.

II

Ι

CAPLUS COPYRIGHT 2006 ACS on STN ANSWER 20 OF 49

ACCESSION NUMBER:

2004:589414 CAPLUS

DOCUMENT NUMBER:

141:134107

TITLE:

A method for the treatment, prevention, or inhibition of a CNS disorder and/or pain and inflammation using a combination of duloxetine, venlafaxine or atomoxetine

and a cyclooxygenase-2 selective inhibitor and

compositions thereof Arneric, Stephen P.

INVENTOR(S): PATENT ASSIGNEE(S):

SOURCE:

Pharmacia Corporation, USA PCT Int. Appl., 208 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004060366	A1	20040722	WO 2003-US38751	20031206

```
AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO,
             NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
             TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
             ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
              TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     US 2004235925
                           A1
                                  20041125
                                             US 2003-727717
                                                                        20031204
     CA 2508884
                           AA
                                  20040722
                                               CA 2003-2508884
                                                                        20031206
     AU 2003294590
                           A1
                                  20040729
                                               AU 2003-294590
                                                                        20031206
     BR 2003017361 ·
                           Α
                                  20051116
                                               BR 2003-17361
                                                                        20031206
PRIORITY APPLN. INFO.:
                                               US 2002-433790P
                                                                     P
                                                                        20021217
                                               WO 2003-US38751
                                                                     W
                                                                        20031206
```

OTHER SOURCE(S): MARPAT 141:134107

A method of treating, preventing, or inhibiting a CNS disorder and/or pain and inflammation or an inflammation-associated disorder in a subject in need of such treatment or prevention provides for treating the subject with duloxetine, venlafaxine or atomoxetine and a cyclooxygenase-2 selective inhibitor or prodrug thereof, wherein the amount of duloxetine, venlafaxine or atomoxetine and the amount of a cyclooxygenase-2 selective inhibitor or prodrug thereof together constitute a CNS disorder, pain and inflammation, or inflammation-associated disorder suppressing treatment, prevention, or inhibition effective amount of the composition

and pharmaceutical compns. that contain duloxetine, venlafaxine or atomoxetine and a cyclooxygenase-2 selective inhibitor are also disclosed.

ANSWER 21 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:589409 CAPLUS

DOCUMENT NUMBER: 141:117197

Compositions and a method for the treatment, TITLE:

prevention, or inhibition of a CNS disorder and/or

pain and inflammation using a combination of

reboxetine and a cyclooxygenase-2 selective inhibitor

INVENTOR(S): Arneric, Stephen P.

PATENT ASSIGNEE(S): Pharmacia Corporation, USA SOURCE: PCT Int. Appl., 192 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT	NO.			KIN	o 1	DATE		į	APPL	ICAT:	ION 1	NO.		D	ATE		
					-									-			
WO 2004	0603	61		A2	:	2004	0722	1	WO 2	003-1	US38'	770		20	00312	205	
WO 2004	0603	51		A3		2004	0902										
W:	ΑE,	AG,	AL,	AM,	ΑT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	ΒZ,	CA,	CH,	
	CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
	GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KΕ,	KG,	ΚP,	KR,	ΚZ,	LC,	
	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NI,	NO,	
	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,	TJ,	
	TM,	TN,	TR,	TT,	TZ,	UA,	ŪĠ,	US,	UΖ,	VC,	VN,	YU,	ZA,	ZM,	ZW		
RW:	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	ΑZ,	
	BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM,	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	
	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	
	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG

```
20031204
    US 2004204411
                      A1
                              20041014 US 2003-727918
                                                                20031205
                        AA
                            '20040722 CA 2003-2510584
    CA 2510584
                      A1
                              20040729 AU 2003-303625
20050921 EP 2003-808444
                                                                20031205
    AU 2003303625
                                                                20031205
                        A2
    EP 1575594
           AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
                                                               20031205
                                        BR 2003-17511
    BR 2003017511
                      Α
                              20051116
                                                                20031205
                        T2
    JP 2006513237
                              20060420
                                          JP 2004-565231
                                                             P 20021217
                                          US 2002-433780P
PRIORITY APPLN. INFO.:
                                          WO 2003-US38770
                                                            W 20031205
```

OTHER SOURCE(S): MARPAT 141:117197

A method of treating, preventing, or inhibiting a CNS disorder and/or pain and inflammation, or an inflammation-associated disorder in a subject in need of such treatment, prevention, or inhibition provides administering to the subject a combination of reboxetine and a cyclooxygenase-2 selective inhibitor or prodrug thereof. Pharmaceutical compns. containing reboxetine and a cyclooxygenase-2 selective inhibitor are also disclosed. For example, a combination of reboxetine and celebrex provided an effective anti-inflammatory activity in a rat carrageenan foot pad edema test, an effective analgesic activity in a rat carrageenan-induced analgesia test, and it was an efficacious treatment for collagen-induced arthritis in mice.

ANSWER 22 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:412933 CAPLUS

DOCUMENT NUMBER:

140:423574

TITLE: Preparation of nitric oxide releasing prodrugs of

diaryl-2-(5H)-furanones as cyclooxygenase-2 inhibitors

INVENTOR(S): Young, Robert N.; Wang, Zhaoyin

PATENT ASSIGNEE(S): Merck Frosst Canada & Co., Can.

SOURCE: PCT Int. Appl., 65 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.						KIND DATE APPLICATION NO.						NO.	DATE					
														-				
	WO 2004041803					A1 20040521			1	WO 2	003-0	CA16		20031103				
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	ΒZ,	CA,	CH,	
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
		GE,	GH,	GM,	HR,	ΗU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KR,	ΚZ,	LC,	LK,	
	•	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NΙ,	NO,	NZ,	
		OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,	ТJ,	TM,	
		TN,	TR,	TT,	TZ,	UA,	ŪĠ,	US,	UΖ,	VC,	VN,	YU,	ZA,	ZM,	ZW			
	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	ΑZ,	BY,	
		KG,	KZ,	MD,	RU,	TJ,	TM,	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	
		FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	TR,	
		BF,	ВJ,	CF,	CG,	CI,	CM,	GA;	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG	
AU 2003283096					A1 20040607				AU 2003-283096						20031103			
PRIORITY APPLN. INFO.:										US 2002-423866P								
									WO 2003-CA1691					W 20031103				

OTHER SOURCE(S): MARPAT 140:423574

GI

$$R^{1}$$
 $O-R^{4}$
 R^{3}
 O
 R^{2}
 I

AB The title compds. I [R1 = SO2Me, etc.; R2, R3 = H, halo, etc.; R4 = NOm, etc.; m = 1 or 2] are prepared The above compds. may be used as a combination therapy with low-dose aspirin to treat chronic cyclooxygenase-2 mediated diseases while simultaneously reducing the risk of thrombotic cardiovascular events.

L8 ANSWER 23 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2004:392439 CAPLUS

DOCUMENT NUMBER:

140:400095

TITLE:

Stereoisomers of p-hydroxy-milnacipran, and

therapeutic use

INVENTOR(S):

Rariy, Roman V.; Heffernan, Michael; Buchwald, Stephen

L.; Swager, Timothy M.

PATENT ASSIGNEE(S):

Collegium Pharmaceutical, Inc., USA

SOURCE:

PCT Int. Appl., 163 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PAT	CENT	NO.			KIN	KIND DATE					ICAT		DATE							
	WO 2004039320 WO 2004039320													20031022						
	W:	AE, CO, GM, LS, PG, TR, GH, KG,	AG, CR, HR, LT, PH, TT, GM, KZ, FR,	AL, CU, HU, LU, PL, TZ, KE, MD, GB,	AM, CZ, ID, LV, PT, UA, LS, RU, GR,	AT, DE, IL, MA, RO, UG, MW, TJ, HU,	AU, DK, IN, MD, RU, UZ, MZ, TM, IE,	AZ, DM, IS, MG, SC, VC, SD, AT, IT,	DZ, JP, MK, SD, VN, SL, BE, LU,	EC, KE, MN, SE, YU, SZ, BG, MC,	EE, KG, MW, SG, ZA, TZ, CH, NL,	ES, KP, MX, SK, ZM, UG, CY, PT,	FI, KR, MZ, SL, ZW, ZM, CZ, RO,	GB, KZ, NI, SY, ZW, DE, SE,	GD, LC, NO, TJ, AM, DK, SI,	GE, LK, NZ, TM, AZ, EE, SK,	GH, LR, OM, TN, BY, ES, TR,			
CA									GN, GQ, GW, ML, MR, NE CA 2003-2503381											
									AU 2003-284342											
	US 2004142904 US 7038085									US 2	003-		20031022							
EP	1578° R:	AT,		CH,	DE,	DK,	2005 ES, RO,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,				
	JP 2006503920 TO PRIORITY APPLN. INFO.:							20060202			JP 2005-501895 US 2002-421640P US 2002-423062P					P 20021025				

US 2003-445142P P 20030205 WO 2003-US33681 W 20031022

OTHER SOURCE(S): MARPAT 140:400095

The invention relates generally to the enantiomers of p-hydroxymilnacipran or congeners thereof. Biol. assays revealed that racemic p-hydroxymilnacipran is approx. equipotent in inhibiting serotonin and norepinephrine uptake (IC50 = 28.6 nM for norepinephrine, IC50 = 21.7 nM for serotonin). Interestingly, (+)-p-hydroxymilnacipran is a more potent inhibitor of norepinephrine uptake than serotonin uptake (IC50 = 10.3 nM for norepinephrine, IC50 = 22 nM for serotonin). In contrast, (-)-p-hydroxymilnacipran is a more potent inhibitor of serotonin uptake compared to norepinephrine uptake (IC50 = 88.5 nM for norepinephrine, IC50 = 40.3 nM for serotonin). The invention also relates to salts and prodrug forms of the above compds. In certain embodiments, the compds. of the invention and a pharmaceutically acceptable excipient are combined to prepare a formulation for administration to a patient. Finally, the invention relates to methods of treating mammals suffering from various afflictions, e.g., depression, chronic pain, or fibromyalgia, comprising administering to a mammal in need thereof a therapeutically effective amount of a compound of the invention. Compound preparation is included.

L8 ANSWER 24 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:101124 CAPLUS

DOCUMENT NUMBER:

140:163574

TITLE:

Preparation of nitric oxide releasing

diaryl-2-(5H)-furanone prodrugs as selective

cyclooxygenase-2 inhibitors for treatment inflammatory

diseases

INVENTOR (S):

Berthelette, Carl; Lachance, Nicholas; Li, Lianhai; Sturino, Claudio; Wang, Zhaoyin; Young, Robert N.;

Dufresne, Claude

PATENT ASSIGNEE(S):

Merck Frosst Canada & Co., Can.

SOURCE:

LANGUAGE:

PCT Int. Appl., 129 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND DATE	APPLICATION NO.				
WO 2004011421	A1 20040205	WO 2003-CA1115	20030724			
W: AE, AG, AL	AM, AT, AU, AZ,	BA, BB, BG, BR, BY, B	Z, CA, CH, CN,			
CO, CR, CU	, CZ, DE, DK, DM,	DZ, EC, EE, ES, FI, G	B, GD, GE, GH,			
GM, HR, HU	, ID, IL, IN, IS,	JP, KE, KG, KR, KZ, LO	C, LK, LR, LS,			
LT, LU, LV	MA, MD, MG, MK,	MN, MW, MX, MZ, NI, NO	O, NZ, OM, PG,			
PH, PL, PT	RO, RU, SC, SD,	SE, SG, SK, SL, SY, To	J, TM, TN, TR,			
TT, TZ, UA	UG, US, UZ, VC,	VN, YU, ZA, ZM, ZW				
RW: GH, GM, KE	LS, MW, MZ, SD,	SL, SZ, TZ, UG, ZM, ZM	W, AM, AZ, BY,			
KG, KZ, MD	RU, TJ, TM, AT,	BE, BG, CH, CY, CZ, DI	E, DK, EE, ES,			
FI, FR, GB	GR, HU, IE, IT,	LU, MC, NL, PT, RO, SI	E, SI, SK, TR,			
BF, BJ, CF	CG, CI, CM, GA,	GN, GQ, GW, ML, MR, NI	E, SN, TD, TG			
CA 2493082	AA 20040205	CA 2003-2493082	20030724			
AU 2003252515	A1 20040216	AU 2003-252515	20030724			
EP 1527045	A1 20050504	EP 2003-771010	20030724			
R: AT, BE, CH	DE, DK, ES, FR,	GB, GR, IT, LI, LU, N	L, SE, MC, PT,			
IE, SI, LT	LV, FI, RO, MK,	CY, AL, TR, BG, CZ, E	E, HU, SK			
US 2005261245	A1 20051124	US 2005-521075	20050112			
PRIORITY APPLN. INFO.:	P 20020726					

US 2002-435341P P 20021220 WO 2003-CA1115 W 20030724

OTHER SOURCE(S): MARPAT 140:163574

GΙ

Title compds. I [R1 = S(0)2CH3, S(0)2NH2, S(0)2NHC(=0)CF3, etc.; R2, R3 =AB H, halo, alkoxy, etc.; R4 = H, (un) substituted alkyl, e.g., halo, Ph, naphthyl, etc.; R5 = NOx, C(=0)-E-alkyl-W-NOx, C(=0)-E-alkyl-Ar-alkyl-W-NOx; x = 1, 2; E = bond, O, S, etc.; W = O, S, C[CO2Rb]2; Ar = CO2Rb(un) substituted Ph, naphthyl, HET3; HET3 = benzimidazolyl, benzofuranyl, benzopyrazolyl, etc.; Rb = (un)substituted alkyl, Ph, naphthyl, etc.] and their pharmaceutically acceptable salts were prepared For example, allylic bromination of Me (2E)-3-[4-(methylsulfonyl)phenyl]-2-phenylbut-2-enoate, e.g., prepared from 1-(4-methanesulfonylphenyl)ethanone in 2 steps, followed by O-alkylation of AqNO3 afforded nitrate ester I [R1 = 4-S(0)2CH3; R2, R3 = H; R4 = CH3; R5 = NO2] in 23% overall yield. In human whole blood LPS induced PGE2 and TXB2 production assays, compds. I have a COX-2 potency and COX-2/COX-1 selectivity comparable to rofecoxib. In paw edema assays in rat, compound I [R1 = 4-S(0)2CH3; R2, R3 = H; R4 = CH3; R5 =CO2(CH2)40NO2] exhibited 42-79% inhibition of pain at 1-30 mg/kg dosage. Of note, compds. I are prodrugs of rofecoxib analogs and are claimed useful for the treatment of chronic COX-2 mediated diseases, while reducing the risk of thrombotic cardiovascular events. Compds. I are useful for treatments of osteoarthritis, rheumatoid arthritis, and chronic pain.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 25 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

Ι

ACCESSION NUMBER: 2004:20448 CAPLUS

DOCUMENT NUMBER: 140:87676

TITLE: Derivatives of gambogic acid and analogs as activators

of caspases and inducers of apoptosis

INVENTOR(S): Tseng, Ben; Sirisoma, Nilantha Sudath; Cai, Sui Xiong;

Zhang, Han-Zhong; Kasibhatla, Shailaja; Ollis, Kristin

P.; Drewe, John A.

PATENT ASSIGNEE(S): Cytovia, Inc., USA

SOURCE: PCT Int. Appl., 92 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

```
KIND
                               DATE
                                          APPLICATION NO.
                                                                 DATE
    PATENT NO.
    -----
                       ----
                               -----
                                          -----
                        A2
                               20040108
                                          WO 2003-US20668
                                                                 20030701
    WO 2004002428
                        A3
                               20050616
    WO 2004002428
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
            PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN,
            TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
            FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                               20040108 CA 2003-2491698
    CA 2491698
                        AA
                                                                 20030701
                                         AU 2003-267977
                                                                 20030701
    AU 2003267977
                         A1
                               20040119
                                        US 2003-609670
EP 2003-748924
                               20040429
    US 2004082066
                         A1
                                                                 20030701
                               20050817
    EP 1562598
                         A2
                                                                 20030701
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
                                          CN 2003-815628
    CN 1738620
                               20060222
                                                                 20030701
                        Α
                         T2
                               20060302
                                          JP 2004-518157
                                                                 20030701
    JP 2006507227
PRIORITY APPLN. INFO.:
                                          US 2002-392358P
                                                            P 20020701
                                          US 2002-413649P
                                                             P 20020926
                                                             W 20030701
                                          WO 2003-US20668
```

OTHER SOURCE(S): MARPAT 140:87676

AB The invention is directed to derivs. of gambogic acid and analogs thereof. Exemplary gambogic acid derivs. of the present invention include, among others, derivs. substituted in the C10 and C28 positions of gambogic acid. The present invention also relates to the discovery that certain preferred compds. of the invention are activators of caspases and inducers of apoptosis. Therefore, the activators of caspases and inducers of apoptosis of this invention can be used to induce cell death in a variety of clin. conditions in which uncontrolled growth and spread of abnormal cells occurs.

L8 ANSWER 26 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:2830 CAPLUS

DOCUMENT NUMBER: 140:59410

TITLE: Preparation of nitrooxy derivatives of

cyclooxygenase-2 inhibitors

INVENTOR(S): Del Soldato, Piero; Santus, Giancarlo

PATENT ASSIGNEE(S): Nicox S.A., Fr.

SOURCE: PCT Int. Appl., 27 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT NO.				D	DATE			APPLICATION NO.						DATE			
WO 2004 WO 2004	A2 20031231 A3 20041014			WO 2003-EP6502						20030620							
	AE, AG CO, CR GM, HR LS, LT	, си, , ни,	AM, CZ, ID,	AT, DE, IL,	AU, DK, IN,	AZ, DM, IS,	DZ, JP,	EC, KE,	EE, KG,	ES, KP,	FI, KR,	GB, KZ,	GD, LC,	GE, LK,	GH, LR,		

```
PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR,
             TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
             FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                            CA 2003-2491209
                                                                    20030620
    CA 2491209
                          AΑ
                                20031231
                                            AU 2003-245972
                                                                    20030620
    AU 2003245972
                          A1
                                20040106
                                            EP 2003-738069
                                                                    20030620
    EP 1517889
                          A2
                                20050330
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
                          Α
                                20050831
                                            CN 2003-814682
                                                                    20030620
    JP 2005530836
                          T2
                                20051013
                                             JP 2004-514803
                                                                    20030620
    ZA 2004010060
                          Α
                                20051020
                                             ZA 2004-10060
                                                                    20041213
    NO 2005000346
                                20050228
                                            NO 2005-346
                                                                    20050121
    US 2006106082
                          A1
                                20060518
                                            US 2005-516938
                                                                    20050913
PRIORITY APPLN. INFO.:
                                             IT 2002-MI1391
                                                                    20020625
                                                                    20030620
                                             WO 2003-EP6502
```

OTHER SOURCE(S): MARPAT 140:59410

Disclosed are new compds. able to release COX-2 inhibitors and NO (no data) having formula M-T-YA-NO2 [wherein M-T = the residue of a COX-2 selective inhibitor (T = SO2NH, SO2NR, CO, O, S, NH, N(SO2R); R = C1-10 alkyl; the COX-2 selective inhibitor, M-TH or M-TOH, has to meet test 2 mentioned in the description); YA = -(B)b0-(C)c0-[b0, c0 = 0,1, with theproviso that b0 and c0 cannot be simultaneously 0; B = TB-X2-TB1; TB = CO, X; X = O, S, NH, NR, R (defined above); TB = CO when T = SO2NH, SO2NR-O,S, NH, or N(SO2R), TB = X when T = CO; TB1 = CO or X (defined above); X2 = a divalent radical selected from the following compds. Q or Q1, etc. (n1, n2 = 0, 1; R2, R3 = H, Me; Y1 = CH2CH2, CH:CH(CH2)n2; n2 = 0, 1)]] for the treatment and/or prophylaxis of inflammatory disorders, pain, fever, cardiovascular disease, gastrointestinal disorders, tumors, Alzheimer's disease, or disorders resulting from elevated levels of COX-2. These compds. including 5-niroxypentanoc acid, 4-nitrooxybutyric acid, and 4-nitrooxybutyramide, 2-nitroxymethylbenzoic acid ester derivs. mitigate or remove the known side-effects of COX-2 inhibitors. The inflammatory disorders are selected from the group consisting of, but not limited to, arthritis, rheumatoid arthritis, osteoarthritis, allergic rhinitis, sinusitis, chronic obstructive pulmonary diseases, dermatitis, psoriasis, cystic fibrosis, multiple sclerosis, vasculitis and organ transplant rejection. The cardiovascular diseases are selected from the group consisting of, but not limited to, atherosclerosis, restenosis, coronary artery disease, angina, diabetes mellitus, diabetic nephropathy, diabetic retinopathy, stroke and myocardial infarct. The qastrointestinal disorders are selected from the group consisting of, but not limited to, inflammatory intestinal disorders, Crohn's disease, qastritis, ulcerative colitis, peptic ulcer, hemorrhagic ulcer, gastric hyperacidity, dyspepsia, gastroparesis, Zollinger-Ellison's syndrome, bacterial infections, hypersecretory states associated with systemic mastocytosis or basophilic leukemia and hyperhystaminemia. The disorders resulting from elevated levels of COX-2 are selected from the group consisting of, but not limited to, angiogenesis, arthritis, asthma, bronchitis, menstrual cramps, tendonitis, bursitis, neoplasia, ophthalmic diseases, pulmonary inflammations, central nervous system disorders, allergic rhinitis, atherosclerosis, endothelial disorders, organs and tissues preservation, inhibition and/or prevention of platelets aggregation. N-[6-[(2,4-difluorophenyl)thio]-2,3-dihydro-1-oxo-1-inden-5-yl]-N-[4-(chloro) butyroyloxymethyl] methanesulfonamide. A solution of chloromethyl (4-chloro) butyrate (1 g, 5.40 mmol) in anhydrous THF (5 mL) was slowly added dropwise in a suspension of N-[6-[(2,4-difluorophenyl)thio]-2,3-dihydro-1oxo-1-inden-5-yl]methanesulfonamide sodium salt (2.04 g, 5.40 mmol) in

anhydrous THF (25 mL) and stirred at room temperature overnight to give, after workup and silica gel chromatog., N-[6-[(2,4-difluorophenyl)thio]-2,3-dihydro-1-oxo-1-inden-5-yl]-N-[4-(chloro)butyroyloxymethyl]methanesulfonam ide (I). A solution of I (1 g, 1.98 mmol) in MeCN (20 mL) was added with AgNO3 (0.67 g, 3.96 mmol), heated at 80° for 15 h in the absence of light, filtered to remove the silver salt, evaporated under vacuum, and purified by chromatog. on a silica gel column to give with n-hexane/ethyl acetate 8/2 as eluent to give 503 mg N-[6-[(2,4-difluorophenyl)thio]-2,3-dihydro-1-oxo-1-inden-5-yl]-N-[4-(nitrooxy)butyroyloxymethyl]methanesulfon amide.

L8 ANSWER 27 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:971878 CAPLUS

DOCUMENT NUMBER: 140:13075

TITLE: Monotherapy for the treatment of amyotrophic lateral

sclerosis with cyclooxygenase-2 (COX 2) inhibitor(s)

INVENTOR(S): Isakson, Peter C.

PATENT ASSIGNEE(S): Pharmacia Corporation, USA SOURCE: PCT Int. Appl., 182 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

```
PATENT NO.
                        KIND
                               DATE
                                         APPLICATION NO.
                                                                 DATE
     _____
                        ----
                               -----
                                           -----
                                                                  -----
    WO 2003101441
                         A1
                               20031211
                                         WO 2003-US14548
                                                                  20030528
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
            PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT,
            TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
            FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
    US 2004063752
                         A1
                               20040401
                                         US 2003-444072
                                                                  20030523
    CA 2487923
                         AA
                               20031211
                                           CA 2003-2487923
                                                                  20030528
    AU 2003232096
                         A1
                               20031219
                                           AU 2003-232096
                                                                  20030528
    BR 2003011518
                               20050222
                                           BR 2003-11518
                                                                  20030528
                         Α
                         A1
                               20050302
                                           EP 2003-756170
                                                                  20030528
    EP 1509217
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
    CN 1658853
                         Α
                               20050824
                                           CN 2003-812637
                                                                  20030528
    JP 2005531592
                         T2
                               20051020
                                           JP 2004-508799
                                                                  20030528
PRIORITY APPLN. INFO.:
                                           US 2002-384139P
                                                               P 20020531
                                           US 2003-444072
                                                               A 20030523
                                           WO 2003-US14548
                                                               W
                                                                  20030528
```

OTHER SOURCE(S): MARPAT 140:13075

AB A method of treating, preventing, or inhibiting amyotrophic lateral sclerosis (ALS), in a subject in need of such treatment, inhibition or prevention. The method comprises administering to a subject one or more cyclooxygenase-2 selective inhibitor(s), or isomer(s), or pharmaceutically acceptable salt(s), ester(s), or prodrug(s) thereof, wherein the amount of the cyclooxygenase-2 selective inhibitor(s), isomer(s), ester(s), salt(s) or prodrug(s) thereof constitutes an ALS treatment, inhibition or prevention effective amount of the COX 2 inhibitor(s).

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS

L8 ANSWER 28 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:971836 CAPLUS

DOCUMENT NUMBER: 140:23256

TITLE: Combination therapy for treatment of amyotrophic

lateral sclerosis (ALS) with cyclooxygenase-2 (COX 2)

inhibitor(s) and a second drug

INVENTOR(S): Isakson, Peter C.

PATENT ASSIGNEE(S): Pharmacia Corporation, USA SOURCE: PCT Int. Appl., 358 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.								APPLICATION NO.									
	2003								1	WO 2	003-1	US14	547		20	0030:	28
WO	2003															1.1.	
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KΕ,	KG,	KP,	KR,	ΚZ,	LC,	LK,	LR,
		LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NI,	NO,	NZ,	OM,
		PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	TJ,	TM,	TN,	TR,	TT,
		TZ,	UA,	ŪĠ,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW					
	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	ŪĠ,	ZM,	ZW,	AM,	ΑZ,	BY,
		KG,	KZ,	MD,	RU,	TJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,
		FI,	FR.	GB,	GR,	HU,	IE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	TR,
		•	•	•	•		CM,					-	-	-	-	-	
us	2004	•	•		•							-	-	-	-	-	
	2487																
	2003																
	2003																
	1539						2005										
EP																	
	R:	•	•	•	•		ES,			•				•			Ρ1,
							RO,										
	2005																
PRIORITY	APP	LN.	INFO	. :													
											003-4		_		A 20		
									. 1	WO 2	003 <i>-</i> 1	JS14	547	ı	v 20	0030	528

OTHER SOURCE(S): MARPAT 140:23256

AB A method of treating, preventing, or inhibiting ALS, in a subject in need of such treatment, inhibition or prevention. The method comprises administering to a subject one or more cyclooxygenase-2 selective inhibitor(s) or isomer(s) or pharmaceutically acceptable salt(s), ester(s), or prodrug(s) thereof, in combination with one or more second drugs, wherein the amount of the cyclooxygenase-2 selective inhibitor(s) or isomer(s) or pharmaceutically acceptable salt(s), ester(s), or prodrug(s) thereof in combination with the amount of second drug(s) constitutes an ALS treatment, inhibition or prevention effective amount

L8 ANSWER 29 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:855795 CAPLUS

DOCUMENT NUMBER: 139:345939

TITLE: Monotherapy for the treatment of Parkinson's disease

with cyclooxygenase 2 (COX2) inhibitor(s)

INVENTOR(S): Stephenson, Diane T.; Isakson, Peter C.; Maziasz,

Timothy J.

PATENT ASSIGNEE(S):

Pharmacia Corporation, USA PCT Int. Appl., 186 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT NO.					KIND DATE			APPLICATION NO.				NO.		DATE			
							-											
	WO	2003	0889	59		A2		2003	1030	1	WO 2	003-1	JS11	517		20	00304	114
	WO	2003	0889	59		A 3		2003	1231									
•		W:	ΑE,	AG,	AL,	AM,	ΑT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
			CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
			GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	ΚP,	KR,	KZ,	LC,	LK,	LR,
												MW,						
			PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	TJ,	TM,	TN,	TR,	TT,
			TZ,	UA,	UG,	US,	UZ,	VC,	VN,	ΥU,	ZA,	ZM,	ZW					
		RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,
			KG,	KZ,	MD,	RU,	TJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,
			FI,	FR,	GB,	GR,	HU,	IE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	TR,
			BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG
	CA	2482	510	•	-	AΑ	·	2003	1030		CA 2	003-	2482	510		20	00304	114
	ΑŲ	2003	2263	79		A1		2003	1103	1	AU 2	003-2	2263	79		20	00304	114
	US	2004	0061	00		A1		2004	0108	1	US 2	003-4	4129	70		20	00304	114
		2003																
	ΕP	1505	962			A2		2005	0216]	EP 2	003-	7469	84		20	00304	114
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
												TR,			-	-	-	•
	JP	2005	•	•	•	•	•	•	•		•			•		•	00304	114
PRIO	RIT	APP	LN.	INFO	. :					1	US 2	002-	3733	17P]	P 20	00204	118
					-							003-					00304	

OTHER SOURCE(S): MARPAT 139:345939

The invention provides a method for treating, preventing, or inhibiting Parkinson's disease (PD), in a subject in need of such treatment, inhibition or prevention. The method comprises treating the subject with one or more COX2 selective inhibitor(s), ester(s), salt(s) or prodrug(s) thereof, wherein the amount of the cyclooxygenase-2 selective inhibitor(s), ester(s), salt(s) or prodrug(s) thereof constitutes a PD treatment-, inhibition- or prevention-effective amount of the COX2 inhibitor(s).

ANSWER 30 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2003:855794 CAPLUS

DOCUMENT NUMBER:

139:345938

TITLE:

Combination therapy including cyclooxygenase 2 (COX2) inhibitor(s) for the treatment of Parkinson's disease

INVENTOR(S):

Stephenson, Diane T.; Isakson, Peter C.; Maziasz, Timothy J.

PATENT ASSIGNEE(S):

Pharmacia Corporation, USA

SOURCE:

PCT Int. Appl., 266 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO.

```
A2
                                20031030
                                           WO 2003-US11269
                                                                   20030414
     WO 2003088958
     WO 2003088958
                         A3
                                20040819
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
            PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT,
            TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
            FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                20031030
                                         CA 2003-2481934
                                                                  20030414
     CA 2481934
                         AΑ
    AU 2003223579
                                20031103
                                          AU 2003-223579
                         A1
                                20040219
                                          US 2003-413348
                                                                  20030414
    US 2004034083
                         A1
    EP 1494664
                         A2
                                20050112
                                           EP 2003-719717
                                                                  20030414
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
                               20050209
                                          BR 2003-9259
                                                                   20030414
    BR 2003009259
                         Α
                         T2
                                20050922
                                           JP 2003-585710
     JP 2005528403
                                                                   20030414
PRIORITY APPLN. INFO.:
                                           US 2002-373311P
                                                                  20020418
                                           WO 2003-US11269
                                                               W 20030414
OTHER SOURCE(S):
                        MARPAT 139:345938
    The invention discloses a method for treating, preventing, or inhibiting
     Parkinson's disease (PD) in a subject in need of such treatment,
     inhibition, or prevention. The method comprises treating the subject with
    one or more COX2 selective inhibitor(s) or isomer(s) or pharmaceutically
     acceptable salt(s), ester(s), or prodrug(s) thereof, in
     combination with one or more second drugs, wherein the amount of the COX2
     selective inhibitor(s) or isomer(s) or pharmaceutically acceptable
     salt(s), ester(s), or prodrug(s) thereof in combination with the
     amount of second drug(s) constitutes a PD treatment-, inhibition- or
    prevention-effective amount
    ANSWER 31 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:
                        2003:656204 CAPLUS
DOCUMENT NUMBER:
                        139:191422
TITLE:
                        Combinations of a cyclooxygenase-2 selective inhibitor
                        and a TNF-\alpha antagonist and therapeutic uses
                        therefor
INVENTOR(S):
                        Bennett, Dennis A.
PATENT ASSIGNEE(S):
                        Pharmacia Corporation, USA
                        U.S. Pat. Appl. Publ., 39 pp.
SOURCE:
                        CODEN: USXXCO
DOCUMENT TYPE:
                        Patent
LANGUAGE:
                        English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
    PATENT NO.
                        KIND
                               DATE
                                          APPLICATION NO.
                                                                  DATE
     ---------
                         ----
                                           ______
                                                                   -----
                                -----
    US 2003157061
                         A1
                               20030821
                                           US 2002-310454
                                                                  20021205
PRIORITY APPLN. INFO.:
                                           US 2001-337802P
                                                              P 20011205
    A method for the prevention, treatment, or inhibition of pain,
    inflammation, or inflammation-related disorder and for the prevention,
    treatment, or inhibition of a cardiovascular disease or disorder in a
    subject that is in need of such prevention, treatment or inhibition,
    involves the administration to the subject of a cyclooxygenase-2 selective
    inhibitor or prodrug thereof and a TNF-\alpha antagonist. A
```

method can also involve the treatment, prevention, or inhibition of cancer

in a subject in need of such treatment, prevention, or inhibition, by administering to the subject a cyclooxygenase-2 selective inhibitor or prodrug thereof and a TNF- α antagonist which is selected from the group consisting of a compound that affects the synthesis of TNF- α , a compound that inhibits the binding of TNF- α with a receptor specific for TNF- α , and a compound that interferes with intracellular signaling triggered by TNF- α binding with a receptor. Compns., pharmaceutical compns. and kits that can be used with the methods are also described.

L8 ANSWER 32 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:633408 CAPLUS

DOCUMENT NUMBER: 139:159977

TITLE: Treatment of colds and cough with a combination of a

cyclooxygenase-2 selective inhibitor and a colds and

cough active ingredient, and compositions thereof

INVENTOR(S): MacMillan, Stephen P.

PATENT ASSIGNEE(S): Pharmacia Corporation, USA

SOURCE: PCT Int. Appl., 147 pp. CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

```
PATENT NO.
                        KIND
                              DATE
                                        APPLICATION NO.
                                                                 DATE
     ______
                        ----
                               -----
                                          _____
                                                                 _____
    WO 2003065988
                        A2
                               20030814
                                          WO 2003-US3221
                                                                 20030204
    WO 2003065988
                        A3
                               20040219
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
            UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
            FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF,
            BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
    CA 2474016
                        AA
                               20030814
                                        CA 2003-2474016
                                                                 20030204
    AU 2003208967
                         A1
                               20030902
                                          AU 2003-208967
                                                                 20030204
                                          US 2003-357747
    US 2004029864
                         A1
                               20040212
                                                                 20030204
                                         EP 2003-707692
    EP 1471872
                        A2
                               20041103
                                                                 20030204
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
    BR 2003007755
                        Α
                               20041207
                                          BR 2003-7755
                                                                 20030204
                         T2
    JP 2005519923
                               20050707
                                           JP 2003-565414
                                                                 20030204
                                          US 2002-354135P
PRIORITY APPLN. INFO.:
                                                              P 20020204
                                          WO 2003-US3221
                                                              W 20030204
```

AB A method for the treatment, prevention and amelioration of colds and/or cough in a subject in need of such treatment, prevention and amelioration, comprises administering to the subject a cyclooxygenase-2 selective inhibitor (e.g. celecoxib; preparation given), or prodrug thereof, and one or more colds and cough active ingredient. Compns., pharmaceutical compns. and kits for practicing the method are also disclosed.

L8 ANSWER 33 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:570770 CAPLUS

DOCUMENT NUMBER: 139:111710

TITLE: Combinations of peroxisome proliferator-activated

receptor- α agonists and cyclooxygenase-2

selective inhibitors, and therapeutic uses therefor

INVENTOR(S): Obukowicz, Mark G.

PATENT ASSIGNEE(S): Pharmacia Corporation, USA SOURCE: PCT Int. Appl., 155 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

		APPLICATION NO.				
WO 2003059294	A2 20030724	WO 2003-US956	20030114			
WO 2003059294	A3 20050714					
W: AE, AG, AL,	AM, AT, AU, AZ,	BA, BB, BG, BR, BY,	BZ, CA, CH, CN,			
CO, CR, CU,	CZ, DE, DK, DM,	DZ, EC, EE, ES, FI,	GB, GD, GE, GH,			
GM, HR, HU,	ID, IL, IN, IS,	JP, KE, KG, KP, KR,	KZ, LC, LK, LR,			
LS, LT, LU,	LV, MA, MD, MG,	MK, MN, MW, MX, MZ,	NO, NZ, OM, PH,			
		SG, SK, SL, TJ, TM,				
UA, UG, US,	UZ, VC, VN, YU,	ZA, ZM, ZW				
RW: GH, GM, KE,	LS, MW, MZ, SD,	SL, SZ, TZ, UG, ZM,	ZW, AM, AZ, BY,			
		BE, BG, CH, CY, CZ,				
FI, FR, GB,	GR, HU, IE, IT,	LU, MC, NL, PT, SE,	SI, SK, TR, BF,			
BJ, CF, CG,	CI, CM, GA, GN,	GQ, GW, ML, MR, NE,	SN, TD, TG			
		US 2003-341217				
		CA 2003-2472168				
AU 2003207535	A1 20030730	AU 2003-207535	20030114			
AU 2003207535						
		JP 2003-559459	20030114			
		EP 2003-705746				
		GB, GR, IT, LI, LU,				
· · · · · · · · · · · · · · · · · · ·		CY, AL, TR, BG, CZ,				
		CN 2003-805942				
		US 2002-348297P				
PRIORITY APPLN. INFO.:		US 2002-340297F				
		WO 2003-341217				
		HO 2003-03936	# 20030114			

OTHER SOURCE(S): MARPAT 139:111710

AB Methods for the treatment, prevention, or inhibition of pain, inflammation, or an inflammation-related disorder, and for the treatment or inhibition of a cardiovascular disease or disorder, and for the treatment or inhibition of cancer, and for the treatment of Alzheimer's disease in a subject in need of such treatment, prevention, or inhibition, include treating the subject with a peroxisome proliferator activated receptor- α agonist and a cyclooxygenase-2 selective inhibitor (e.g. celecoxib; preparation described), or prodrug thereof. Compns., pharmaceutical compns., and kits for effecting the particular methods are also described.

L8 ANSWER 34 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:570750 CAPLUS

DOCUMENT NUMBER: 139:111706

TITLE: peroxisome proliferator-activated receptor-α

agonist- and cyclooxygenase-2 selective

inhibitor-containing compositions, and methods of

treatment using them

INVENTOR(S): Needleman, Philip

PATENT ASSIGNEE(S): Pharmacia Corporation, USA SOURCE: PCT Int. Appl., 157 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

						KIND DATE			APPLICATION NO.						DATE			
											2003-t							
WO	2003	0592	71		A3	•	2003	1127										
	W:									BB	, BG,	BR,	BY,	BZ,	CA,	CH,	CN,	
		co,	CR.	CU,	cz,	DE,	DK,	DM,	DZ,	EC	, EE,	ES,	FI,	GB,	GD,	GE,	GH,	
		GM.	HR.	HU,	ID,	IL.	IN,	ıs,	JP,	KE	, KG,	KP,	KR,	KZ,	LC,	LK,	LR,	
											, MW,							
											, SL,							
											, ZW	•	•	•	•		•	
	RW:	-									, TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,	
											, сн,							
											, NL,							
											, ML,							
us	2003										2003-:						113	
											2003-2							
											2003-2							
	2003																	
EP	1465	621			A2		2004	1013		EP :	2003-	7057	68		2	0030	114	
											, IT,							
											, TR,						•	
CN	1642		•	•	A	•	2005	0720		CN :	2003-	8058	86	•	2	0030	114	
BR	2003	0068	72		Α		2005	0906		BR :	2003-0	6872			2	0030	114	
JP	2006	5011	36		Т2		2006	0112	1	JP :	2003-	5594	36		2	0030	114	
											2004-					0040		
PRIORIT										US :	2002-	3482	98P		P 2	0020	114	
	_ .			-							2003-					0030		
											2003-1					0030	114	
		/a\				~ ~ ~					_							

OTHER SOURCE(S): MARPAT 139:111706

AB Methods for the treatment, prevention, or inhibition of pain, inflammation, or inflammation-related disorder, and for the treatment or inhibition of a cardiovascular disease or disorder, and for the treatment or inhibition of cancer in a subject in need of such treatment, prevention, or inhibition, include treating the subject with a peroxisome proliferator activated receptor-α agonist and a cyclooxygenase-2 selective inhibitor (e.g. celecoxib; preparation described), or prodrug thereof. Compns., pharmaceutical compns., and kits for effecting the particular methods are also described.

L8 ANSWER 35 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:551339 CAPLUS

DOCUMENT NUMBER: 139:95464

TITLE: Treatment of pain, inflammation, and

inflammation-related disorders with a combination of a

cyclooxygenase-2 selective inhibitor and aspirin

INVENTOR(S): Macmillan, Stephen P.

PATENT ASSIGNEE(S): Pharmacia Corporation, USA SOURCE: PCT Int. Appl., 123 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

```
KIND DATE APPLICATION NO.
                                                             DATE
                     ----
                                       _____
    -----
                                                             _____
    WO 2003057166 A2 20030717 WO 2003-US255 WO 2003057166 A3 20031106
                                                              20030107
           AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
           CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
           GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
           LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
           PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
           UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
           KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
           FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF,
           BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                       AA 20030717 CA 2003-2471951
    CA 2471951
    AU 2003207453
                       A1
                             20030724
                                      AU 2003-207453
                                                             20030107
                      A2
    AU 2003207453
                             20030724
                                                             20030107
    US 2003143271
                      A1
                            20030731 US 2003-337583
    US 2003217846
                      A1 20031106 US 2003-337760
A2 20041027 EP 2003-705660
                      A1
    EP 1469846
                                                              20030107
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
           IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
    BR 2003006777 A 20050426 BR 2003-6777
                                       CN 2003-805349
    CN 1638760
                       Α
                             20050713
                                                             20030107
                                      JP 2003-557525
    JP 2005524618
                       T2
                             20050818
                                      ZA 2004-5379
    ZA 2004005379
                      A 20050617
                                       US 2002-346560P P 20020107
WO 2003-US255 W 20030107
PRIORITY APPLN. INFO.:
```

AB A method for the prevention, treatment, or amelioration of pain, inflammation, or inflammation-related disorder in a subject that is in need of such prevention, treatment or amelioration, involves the administration to the subject of a cyclooxygenase-2 selective inhibitor or prodrug thereof and enteric-coated aspirin. A method can also involve the administration of a cyclooxygenase-2 selective inhibitor and aspirin in an amount lower than 75 mg/day. A method can also involve the administration of a cyclooxygenase-2 selective inhibitor and aspirin where the cyclooxygenase-2 selective inhibitor is BMS-347070, S-33516, CS-502, darbufelone, LAS 34475, LAS 34556, L-745337, SD-8381, RWJ-63556, L-784512, COX-189, ABT-963, or valdecoxib, or any pharmaceutical salt or prodrug thereof. Compns., pharmaceutical compns., and kits that can be used with the methods are also described. Preparation of celecoxib is described.

```
L8 ANSWER 36 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN
```

ACCESSION NUMBER: 2003:532347 CAPLUS

DOCUMENT NUMBER: 139:79173

TITLE: Methods and compositions using a cyclooxygenase 2

(COX-2) inhibitor for the treatment of psychiatric

disorders

INVENTOR(S):
Muller, Norbert

PATENT ASSIGNEE(S): Germany

SOURCE: U.S. Pat. Appl. Publ., 27 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

```
20030710
                                            US 2002-157969
    US 2003130334
                          A1
                                                                   20020531
                                            EP 2005-24864
     EP 1627639
                          A2
                                20060222
                                                                   20020531
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
                                20060727
                                            US 2005-320757
     US 2006167074
                         A1
                                                                    20051230
PRIORITY APPLN. INFO.:
                                            DE 2001-10129328
                                                                A 20010619
                                            US 2002-364904P
                                                                P 20020314
                                            DE 2001-10129320
                                                                A 20010619
                                            EP 2002-738138
                                                                A3 20020531
                                            US 2002-157969
                                                                A2 20020531
OTHER SOURCE(S):
                         MARPAT 139:79173
```

AB A method for the prevention, treatment, or inhibition of a psychiatric disorder, in particular schizophrenia, is described which comprises administering a COX-2 inhibitor, or prodrug thereof, to a subject. Moreover, a method for the prevention, treatment, or inhibition of a psychiatric disorder, in particular schizophrenia or a depressive disorder, is disclosed, comprising administering to a subject a COX-2 inhibitor or prodrug thereof in combination with a neuroleptic drug or an antidepressant. Compns. and kits that are suitable for the practice of the method are also described.

L8 ANSWER 37 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:492716 CAPLUS

DOCUMENT NUMBER: 139:63316

TITLE: Methods using a combination of a 3-heteroaryl-2-

indolinone and a cyclooxygenase-2 inhibitor for the

treatment of neoplasia

INVENTOR(S): Masferrer, Jaime L.; Cherrington, Julie M.; Leahy,

Kathleen M.; Zweifel, Ben S.

PATENT ASSIGNEE(S): Pharmacia Corporation, USA

SOURCE: U.S. Pat. Appl. Publ., 66 pp., Cont.-in-part of Appl.

No. PCT/US99/30693.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 21

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.				
US 2003119895			20020516			
WO 2000038730	A3 20001102					
W: AE, AL, AM,	AT, AU, AZ, BA,	BB, BG, BR, BY, CA, CH,	CN, CR, CU,			
CZ, DE, DK,	DM, EE, ES, FI,	GB, GD, GE, GH, GM, HR,	HU, ID, IL,			
IN, IS, JP,	KE, KG, KP, KR,	KZ, LC, LK, LR, LS, LT,	LU, LV, MA,			
MD, MG, MK,	MN, MW, MX, NO,	NZ, PL, PT, RO, RU, SD,	SE, SG, SI,			
SK, SL, TJ,	TM, TR, TT, TZ,	UA, UG, US, UZ, VN, YU,	ZA, ZW			
RW: GH, GM, KE,	LS, MW, SD, SL,	SZ, TZ, UG, ZW, AT, BE,	CH, CY, DE,			
DK, ES, FI,	FR, GB, GR, IE,	IT, LU, MC, NL, PT, SE,	BF, BJ, CF,			
CG, CI, CM,	GA, GN, GW, ML,	MR, NE, SN, TD, TG				
EP 1522313	A1 20050413	EP 2004-26577	19991222			
R: AT, BE, CH,	DE, DK, ES, FR,	GB, GR, IT, LI, LU, NL,	SE, MC, PT,			
IE, FI, RO,	CY					
CA 2484324	AA 20031127	CA 2003-2484324	20030515			
WO 2003097044	A1 20031127	WO 2003-US15582	20030515			
W: AE, AG, AL,	AM, AT, AU, AZ,	BA, BB, BG, BR, BY, BZ,	CA, CH, CN,			
CO, CR, CU,	CZ, DE, DK, DM,	DZ, EC, EE, ES, FI, GB,	GD, GE, GH,			
GM, HR, HU,	ID, IL, IN, IS,	JP, KE, KG, KP, KR, KZ,	LC, LK, LR,			
LS, LT, LU,	LV, MA, MD, MG,	MK, MN, MW, MX, MZ, NI,	NO, NZ, OM,			

```
PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT,
            TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
            FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                        AU 2003-239494
     AU 2003239494
                         A1
                               20031202
                                                                 20030515
                                         BR 2003-10027
                               20050215
                                                                 20030515
     BR 2003010027
                         Α
                                         EP 2003-734058
                               20050302
                                                                 20030515
     EP 1509224
                         A1
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     JP 2005530781
                        T2
                               20051013
                                          JP 2004-505043
                                                                  20030515
     AU 2004210578
                         A1
                               20041007
                                           AU 2004-210578
                                                                  20040910
PRIORITY APPLN. INFO.:
                                           US 1998-113786P
                                                             P 19981223
                                           WO 1999-US30693
                                                             A2 19991222
                                           US 1999-385214
                                                              A 19990827
                                           AU 2000-25936
                                                              A3 19991222
                                           EP 1999-968939
                                                              A3 19991222
                                           US 2002-150546
                                                              A 20020516
                                                               W 20030515
                                           WO 2003-US15582
                        MARPAT 139:63316
OTHER SOURCE(S):
AB
     The invention provides methods and compns. useful for treatment or
     prevention of neoplasia by administering a combination comprising a
     3-heteroaryl-2-indolinone compound (preparation included) and a COX-2 selective
     inhibitor. Further provided are compns., pharmaceutical compns., and kits
     for treatment and prevention of neoplasia.
    ANSWER 38 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:
                        2003:154262 CAPLUS
DOCUMENT NUMBER:
                        138:198610
                        Compositions for the treatment and prevention of pain
TITLE:
                        and inflammation with a cyclooxygenase-2 selective
                        inhibitor and chondroitin sulfate
INVENTOR (S):
                        Pulaski, Steven P.; Kundel, Susan
PATENT ASSIGNEE(S):
                        Pharmacia Corporation, USA
SOURCE:
                        PCT Int. Appl., 148 pp.
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
LANGUAGE:
                        English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                                         APPLICATION NO.
    PATENT NO.
                        KIND
                               DATE
                                                                 DATE
     -----
                        ----
                               -----
                                          -----
                                                                 _____
                                        WO 2002-US25673
    WO 2003015799
                        A1
                               20030227
                                                                20020813
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
```

```
GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
        LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
        PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
        UA, UG, US, UZ, VN, YU, ZA, ZM, ZW
    RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
        CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
        PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
        NE, SN, TD, TG
US 2003114416
                     A1
                           20030619
                                       US 2002-215539
                                                               20020809
CA 2457452
                     AA
                           20030227
                                       CA 2002-2457452
                                                              20020813
AU 2002336344
                     A2
                           20030303
                                       AU 2002-336344
                                                              20020813
EP 1416941
                    A1
                           20040512
                                      EP 2002-773188
                                                              20020813
    R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
```

```
IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK
                                                                  20020813
     BR 2002011977
                         Α
                               20040921
                                        BR 2002-11977
                         T2
                                                                  20020813
     JP 2005501850
                               20050120
                                           JP 2003-520758
                         Α
                               20050202
                                           CN 2002-820121
                                                                  20020813
    CN 1575182
                         Α
                               20050622
                                           ZA 2004-1163
                                                                  20040212
     ZA 2004001163
                                                               P 20010814
PRIORITY APPLN. INFO.:
                                           US 2001-312211P
                                           US 2002-215539
                                                               A 20020809
                                           WO 2002-US25673
                                                               W
                                                                  20020813
                        MARPAT 138:198610
OTHER SOURCE(S):
    A method of treating, preventing, or inhibiting pain, inflammation, or
     inflammation-associated disorder in a subject in need of such treatment or
    prevention includes treating the subject with chondroitin sulfate and a
    cyclooxygenase-2 selective inhibitor, or a prodrug thereof,
    wherein the amount of chondroitin sulfate and the amount of a cyclooxygenase-2
     selective inhibitor or a pharmaceutically acceptable salt or
    prodrug thereof together constitute a pain- or
     inflammation-suppressing treatment or prevention effective amount
    Glucosamine can optionally be present. Compns. that contain the
     combination of chondroitin sulfate and cyclooxygenase-2 selective
     inhibitor and, optionally, the glucosamine, are disclosed, as are
    pharmaceutical compns.
REFERENCE COUNT:
                              THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS
                              RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 39 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:
                        2003:154260 CAPLUS
DOCUMENT NUMBER:
                        138:198609
TITLE:
                        Compositions for the treatment and prevention of pain
                        and inflammation with a cyclooxygenase-2 selective
                        inhibitor and glucosamine
INVENTOR(S):
                        Pulaski, Steven P.; Kundel, Susan
                        Pharmacia Corporation, USA
PATENT ASSIGNEE(S):
                        PCT Int. Appl., 145 pp.
SOURCE:
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
                        English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
    PATENT NO.
                        KIND
                               DATE
                                         APPLICATION NO.
                                                                  DATE
     ______
                        ____
                               -----
                                           -----
                                                                  _____
    WO 2003015797
                        A1
                               20030227
                                          WO 2002-US25674
                                                                  20020813
    WO 2003015797
                        C1
                               20041229
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
            UA, UG, US, UZ, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
            CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
            PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
            NE, SN, TD, TG
    US 2003114418
                         A1
                               20030619
                                           US 2002-215816
                                                                  20020809
    CA 2457453
                         AA
                               20030227
                                           CA 2002-2457453
                                                                  20020813
    AU 2002331076
                         A2
                               20030303
                                         AU 2002-331076
                                                                  20020813
                                          EP 2002-768522
    EP 1416940
                         A1
                               20040512
                                                                  20020813
```

AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

BR 2002-11936

20020813

IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK

20041026

Α

```
20020813
    JP 2005507871
                         T2
                               20050324
                                          JP 2003-520756
                                          CN 2002-820216
                                                                 20020813
                               20060503
    CN 1767835
                         Α
                                                                 20040212
                         Α
                               20050622
                                          ZA 2004-1158
    ZA 2004001158
                                          US 2001-312272P
                                                              P
                                                                 20010814
PRIORITY APPLN. INFO.:
                                                              Α
                                                                 20020809
                                          US 2002-215216
                                                                 20020809
                                                              Δ
                                          US 2002-215816
                                          WO 2002-US25674
                                                                 20020813
```

OTHER SOURCE(S): MARPAT 138:198609

A method of treating, preventing, or inhibiting pain, inflammation or inflammation-associated disorder in a subject in need of such treatment or prevention provides for treating the subject with glucosamine and a cyclooxygenase-2 selective inhibitor or prodrug thereof, wherein the amount of glucosamine and the amount of a cyclooxygenase-2 selective inhibitor or prodrug thereof together constitute a pain or inflammation suppressing treatment or prevention effective amount of the composition Compns. and pharmaceutical compns. that contain glucosamine and a cyclooxygenase-2 selective inhibitor are also disclosed.

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 4 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 40 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:154230 CAPLUS

DOCUMENT NUMBER: 138:210277

Synthesis and use of reagents for improved DNA TITLE:

lipofection and/or slow release prodrug and

drug therapies

INVENTOR(S): Diamond, Scott L.; Gruneich, Jeffrey

PATENT ASSIGNEE(S): The Trustees of the University of Pennsylvania, USA

SOURCE: PCT Int. Appl., 70 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION: DAMENTO NO

	PATENT NO.						APPLICATION NO.										
	2003														2	0020	315
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
		co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,
		•	•	•	•						MW,	•					
			•	•	•						SL,			•			-
				•	UZ,			-			•	•		•		•	•
	RW:	•	•	•	•						TZ,	UG.	ZM.	ZW.	AT.	BE.	BG.
	••••										GB,						
		•	•	•	•						CM,	•				-	-
		•	SN,	•	•	22,	20,	 ,	υυ,	O-,	J,	J.,	U 11,	- 2,	J,	,	,
CA	2456	•	•	•			2003	0227	(כ בי	002-	2456	977		2	0020	815
	1424															0020	
BF											IT,						
	к.			-							TR,					110,	,
TD	2005			•	•	-	-			-	1003-		-			0020	015
	2005				AI		2005	0331									
PRIORIT	Y APP	LN.	INFO	. :							001-						
											002-					0020	
አ ው ሞኤ	o inte			. 1							002-1				_		

The invention relates to compns. and methods for a one-step synthetic AB technique for making cationic steroid or cationic drug mols. for use as delivery vehicles. The invention further relates to methods for using

Page 49

cationic steroid mols. in lipofection or transfection, delivery of drugs, and for treatment of inflamrnation and other diseases and disorders. The invention also relates to cationic steroid prodrugs and cationic prodrugs and to methods of modifying drugs.

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS 4 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 41 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2002:977588 CAPLUS

DOCUMENT NUMBER:

138:33362

TITLE:

Use of cyclooxygenase 2 (COX-2) inhibitors for the treatment of schizophrenia, delusional disorders, affective disorders, autism, or tic disorders

INVENTOR(S):

Muller, Norbert

PATENT ASSIGNEE(S):

Germany

SOURCE:

PCT Int. Appl., 58 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION: DAMENTO NO

	PATENT NO.				KIND DATE		APPLICATION NO.						DATE 20020531					
																_	:	
										1	WO 2	002-	EP60	13		2	0020	531
1	WO	2002																
		W:	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	ВG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
			CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FΙ,	GB,	GD,	GE,	GH,
			GM,	HR,	ΗU,	ID,	IL,	IN,	IS,	JΡ,	KΕ,	KG,	ΚP,	KR,	KZ,	LC,	LK,	LR,
			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	OM,	PH,
			PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TN,	TR,	TT,	TZ,
			UA,	UG,	US,	UZ,	VN,	YU,	ZA,	ZM,	ZW			•				
		RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,
			KG,	KZ,	MD,	RU,	TJ,	TM,	AT,	BE,	CH,	CY,	DE,	DK,	ES,	FI,	FR,	GB,
								NL,										
								NE,					•	_	-			
	DE	1012						2003				001-	1012	9320		2	0010	619
	CA	2448	025															
		1397																
		1397																
•								ES,		GB.	GR.	IT.	LI.	LU.	NL.	SE.	MC.	PT.
		20.				•	•	RO,	•		•	•	,	,	,	,	,	,
	TP.	2004	5340	66	,	Т2	,	2004	1111	U-,	TP 2	003-	5048	86		2	0020	531
		1627																
								ES,										
		10.						RO,					,	_0,	112,	,	,	~ - ,
	110	2004											4806	0.0		2	0040	205
PRIOR						A.		2001									0010	
PRIOR	111	APP.	TITA .	INFO	• •									04P		_	0020	
														38				
														13			0020	
OTHER	sc	URCE	(S):			MARI	PAT	138:	3336		NO 2	002-	EFQU.	13	,	,, 2	0020	J J I

The invention discloses the use of a COX-2 inhibitor for the treatment of psychiatric disorders, e.g. schizophrenia, delusional disorders, affective disorders, autism or tic disorders, in particular chronic schizophrenic psychoses and schizoaffective psychoses, temporary acute psychotic disorders, depressive episodes, recurring depressive episodes, manic episodes and bipolar affective disorders. Moreover, the invention discloses the use of a COX-2 inhibitor, in particular celecoxib, in combination with a neuroleptic drug, in particular risperidone, or an

antidepressant, for the treatment of psychiatric disorders such as schizophrenia, delusional disorders, affective disorders, autism or tic disorders.

L8 ANSWER 42 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:927258 CAPLUS

DOCUMENT NUMBER: 138:16609

TITLE: Skin-permeable selective cyclooxygenase-2 inhibitor

composition

INVENTOR(S): Lu, Guang Wei; Ewing, Gary D.; Tyle, Praveen; Stoller,

Brenda M.; Gokhale, Rajeev; Gadre, Ashwini

PATENT ASSIGNEE(S): Pharmacia Corporation, USA

SOURCE: PCT Int. Appl., 55 pp.

CODEN: PIXXD2
DOCUMENT TYPE: Patent

LANGUAGE: Patent English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

P					KIND DATE			APPLICATION NO.				DATE						
								2002		,	WO	2002-1	US17	067		2	0020	530
		W:										, BG,						
												, KG,						
			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN	, MW,	MX,	MZ,	NO,	ΝZ,	OM,	PH,
			PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK	, SL,	ΤJ,	TM,	TN,	TR,	TT,	TZ,
			UA,	UG,	US,	UΖ,	VN,	YU,	ZA,	ZM,	ZW							
		RW:	GH,	GM,	ΚE,	LS,	MW,	MZ,	SD,	SL,	SZ	, TZ,	ΨG,	ZM,	ZW,	AM,	ΑZ,	BY,
			KG,	ΚZ,	MD,	RU,	TJ,	TM,	ΑT,	BE,	CH	CY,	DE,	DK,	ES,	FI,	FR,	GB,
			GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	TR	, BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,
		•	GN,		-	-		ΝE,										
C	Ά	24486	527			AA		2002	1205	(CA	2002-	2448	527		2	0020	530
												2002-						
												2002-						
E	P	14043	345			A2		2004	0407	:	ΕP	2002-	7741	23		2	0020	530
		R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR	, IT,	LI,	LU,	NL,	SE,	MC,	PT,
			ΙE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL	, TR						
В	R	20020	0101	04		Α		2004	0608]	BR	2002-	10104	4		2	0020	530
J	Ρ	20045	5328	71		T2		2004				2002-					0020	530
C	N	15474	174			Α		2004	1117	(CN	2002-	81494	46		2	0020	530
Z	Α	20030	0092	98		Α		2004	0512		ZA	2003-	9298			2	0031	128
PRIORI	TY	APPI	١N. :	INFO	. :					1	US	2001-	2948	38P]	P 2	0010	531
										1	US	2001-	3507	56P]	P 2	0011	113
										1	WO	2002-1	US17	067	1	1 2	0020	530
OTHER	SO	URCE	(s)			MARI	РΑТ	138	16609	9								

OTHER SOURCE(S): MARPAT 138:16609

AB A skin deliverable pharmaceutical composition comprises at least 1 selective cyclooxygenase-2 (COX-2) inhibitory drug or prodrug thereof solubilized in a pharmaceutically acceptable carrier that contains a low mol. weight monohydric alc., and exhibits a skin permeation rate of the therapeutic agent at least equal to that exhibited by a reference solution of the

therapeutic agent in 70% aqueous ethanol. A method of effecting targeted delivery of a selective COX-2 inhibitory drug to a site of pain and/or inflammation in a subject comprises topically administering such a composition to skin of the subject, preferably at a locus overlying or adjacent to the site of pain and/or inflammation. A method of effecting systemic treatment of a subject having a COX-2 mediated disorder comprises transdermally administering such a composition, preferably by contacting the

Page 51

composition with an area of skin of the subject ≥400 cm2. Thus, celecoxib was observed in 70% aqueous EtOH and this solution provided greater skin

flux of the drug.

L8 ANSWER 43 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:556104 CAPLUS

DOCUMENT NUMBER: 137:109489

TITLE: Compositions comprising a polypeptide and an active

agent

INVENTOR(S): Piccariello, Thomas; Olon, Lawrence P.; Kirk, Randal

J.

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 34 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 20

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
US 2002099013	A1	20020725	US 2001-933708		20010822
US 2004087483	A1	20040506	US 2002-136433		20020502
US 2004063628	A1	20040401	US 2002-156527		20020529
US 7060708	B2	20060613			
US 2006014697	A1	20060119	US 2005-89056		20050325
PRIORITY APPLN. INFO.:			US 2000-247556P	P	20001114
			US 2000-247558P	P	20001114
			US 2000-247559P	P	20001114
			US 2000-247560P	P	20001114
			US 2000-247561P	P	20001114
			US 2000-247594P	P	20001114
			US 2000-247595P	P	20001114
			US 2000-247606P	P	20001114
			US 2000-247607P	P	20001114
			US 2000-247608P	P	20001114
			US 2000-247609P	P	20001114
			US 2000-247610P	P	20001114
			US 2000-247611P	P	20001114
			US 2000-247612P	P	20001114
			US 2000-247620P	P	20001114
			US 2000-247621P	P	20001114
			US 2000-247634P	P	20001114
			US 2000-247635P	P	20001114
			US 2000-247698P	P	20001114
			US 2000-247699P	P	20001114
			US 2000-247700P	P	20001114
			US 2000-247701P	P	20001114
			US 2000-247702P	P	20001114
			US 2000-247797P	P	20001114
			US 2000-247798P	P	20001114
			US 2000-247799P	P	20001114
			US 2000-247800P	P	20001114
			US 2000-247801P	P	20001114
			US 2000-247802P	P	20001114
,			US 2000-247803P	P	20001114
			US 2000-247804P	P	20001114
			US 2000-247805P	P	20001114
				_	

US 2000-247807P P 20001114

US	2000-247832P	P	20001114
US	2000-247833P	Р	20001114
		_	
US	2000-247926P	P	20001114
US	2000-247927P	P	20001114
US	2000-247928P	P	20001114
US	2000-247929P	P	20001114
US	2000-247930P	P	20001114
US	1999-265415	В2	19990310
US	1999-411238	B2	19991004
WO	2000-US5693	Α	20000306
US	2000-642820	A2	20000822
US	2000-248607P	P	20001116
US	2000-248620P	P	20001116
US	2000-248656P	P	20001116
US	2000-248658P	P	20001116
US	2000-248659P	P	20001116
US	2000-248660P	P	20001116
US	2000-248662P	P	20001116
US	2000-248663P	P	20001116
US	2000-248685P	P	20001116
US	2000-248737P	P	20001116
US	2000-248738P	P	20001116
US	2000-248764P	P	20001116
US	2000-248767P	P	20001116
US	2000-248768P	P	20001116
US	2000-248769P	P	20001116
US	2000-248770P	P	20001116
US	2000-248771P		20001116
		P	
US	2000-248772P	P	20001116
US	2000-248774P	P	20001116
US	2000-248776P	P	20001116
US	2000-248777P	P	20001116
US	2000-248778P	P	20001116
US	2000-248779P	P	20001116
		_	
US	2000-248782P	P	20001116
US	2000-248787P	P	20001116
US	2000-248794P	P	20001116
US	2000-248795P	P	20001116
US	2000-248796P	P	20001116
US	2000-248797P	P	20001116
US	2001-933708	A2	20010822
US	2001-986426	A2	20011108
US	2001-987458	B2	20011114
WO	2001-US43089	B2	20011114
US	2001-988034	B2	20011116
US	2001-988071	B2	20011116
WO	2001-US43115	B2	20011116
WO	2001-US43117	B2	20011116
US	2002-358368P	P	20020222
US	2002-358381P	P	20020222
US	2002-362082P	P	20020307
US	2002-366258P	P	20020322
	2002-3662387		20020522
US		A2	
WO	2003-US5525	A2	20030224
US	2003-507012P	P	20030930
US	2004-567800P	P	20040505
US	2004-567802P	P	20040505
US	2004-568011P	P	20040505
US	2004 - 923088	A2	20040823
US	2004-923257	A2	20040823

US 2004-953110 A2 20040930 US 2004-953111 A2 20040930 US 2004-953116 A2 20040930 US 2004-953119 A2 20040930 US 2004-955006 A2 20040930 WO 2004-US32131 A2 20040930

AB Claimed are compns. comprising a polypeptide and an active agent covalently attached to the polypeptide and a method for delivery of an active agent to a patient by administering the composition to the patient. The peptide is a homopolymer of a naturally occurring amino acid or a heteropolymer of two or more naturally occurring amino acids. In an example, (Glu)n-cephalexin was prepared from Glu(OBut)NCA and cephalexin hydrochloride.

L8 ANSWER 44 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:392237 CAPLUS

DOCUMENT NUMBER: 136:401651

TITLE: Preparation of fused pyridine derivatives as HMG-COA

reductase inhibitors

INVENTOR(S): Robl, Jeffrey A.; Chen, Bang-Chi; Sun, Chong-Qing

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 46 pp., Cont.-in-part of U.S.

Ser. No. 875,218. CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002061901	A1	20020523	US 2001-8154	20011204
US 6620821	B2	20030916		
US 2002028826	A1	20020307	US 2001-875218	20010606
US 2004024216	A1	20040205	US 2003-602753	20030624
PRIORITY APPLN. INFO.:			US 2000-211594P P	20000615
			US 2001-875218 A	2 20010606
			US 2001-8154 A	3 20011204

OTHER SOURCE(S): MARPAT 136:401651

GI

$$R^2$$
 R^2
 $N = (O)_n$
 CO_2Na
 R^4
 R^4
 R^4
 R^4
 R^4
 R^4
 R^4

The title compds. I and their pharmaceutically acceptable salts, esters, AB prodrug esters, and stereoisomers are claimed [wherein: Z = CH(OH)CH2CR7(OH)CH2CO2R3 or corresponding pyranone lactone derivs.; n = 0, 1; x = 0, 1, 2, 3, or 4; y = 0, 1, 2, 3 or 4, provided that at least one of x and y is other than 0; and optionally one or more carbons of (CH2)x and/or (CH2)y together with addnl. carbons form a 3 to 7 membered spirocyclic ring; R1, R2 = alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl, cycloheteroalkyl; R3 = H or lower alkyl; R4 = H, halo, CF3, OH, alkyl, alkoxy, CO2H, (un) substituted NH2, cyano, (un) substituted CONH2, etc.; R7 = H, alkyl]. The compds. are HMG-CoA reductase inhibitors, and are active in inhibiting cholesterol biosynthesis and modulating blood serum lipids, for example, lowering LDL cholesterol and/or increasing HDL cholesterol (no data). I are thus useful in treating hyperlipidemia and dyslipidemia, in hormone replacement therapy, and in treating hypercholesterolemia, hypertriglyceridemia and atherosclerosis, as well as Alzheimer's disease and osteoporosis. Prepns. of several compds. are described. For instance, a multistep synthesis of fused pyridine derivative II is reported. Compds. I may be used in a manner similar to atorvastatin, pravastatin, simvastatin, etc. Combinations of compds. I with various other drugs are claimed, the latter being specified as certain pharmacol. classes, as inhibitors of specific enzymes, as (ant) agonists of specific receptors, and as numerous named drugs.

L8 ANSWER 45 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:332011 CAPLUS

DOCUMENT NUMBER: 136:355482

TITLE: Compositions comprising a polypeptide and an active

agent

INVENTOR(S): Piccariello, Thomas; Olon, Lawrence P.; Kirk, Randall

J.

PATENT ASSIGNEE(S): New River Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 98 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 20

PATENT INFORMATION:

PATENT NO.		APPLICATION NO.			
WO 2002034237	A1 20020502	WO 2001-US26142	20010822		
W: AE, AG, AL	, AM, AT, AU, AZ,	BA, BB, BG, BR, BY, BZ,	CA, CH, CN,		
CR, CU, CZ	, DE, DK, DM, DZ,	EE, ES, FI, GB, GD, GE,	GH, GM, HR,		
HU, ID, IL	, IN, IS, JP, KE,	KG, KP, KR, KZ, LC, LK,	LR, LS, LT,		
• •		MW, MX, MZ, NO, NZ, PL,			
•		TM, TR, TT, TZ, UA, UG,			
ZA, ZW	, 51, 51, 52, 10,	11., 12, 11, 12, 011, 00,	02, 111, 20,		
•	TO MIL MY CD	CI C7 T7 IIC 7W AT	DE CU'CV		
• •		SL, SZ, TZ, UG, ZW, AT,			
•		IE, IT, LU, MC, NL, PT,			
		GQ, GW, ML, MR, NE, SN,	•		
US 6716452	B1 20040406	US 2000-642820	20000822		
CA 2420590	AA 20020502	CA 2001-2420590	20010822		
AU 2001086599	A5 20020506	AU 2001-86599	20010822		
EP 1311242	A1 20030521	EP 2001-966056	20010822		
		GB, GR, IT, LI, LU, NL,			
	, LV, FI, RO, MK,				
		JP 2002-537291	20010822		
<u>-</u>		US 2003-727565			
PRIORITY APPLN. INFO.:		US 2000-642820	A 20000822		

```
20001114
US 2000-247613P
                     Ρ
                        20001114
US 2000-247614P
                     Р
US 2000-247615P
                     Р
                        20001114
US 2000-247616P
                     Ρ
                        20001114
US 2000-247617P
                     Р
                        20001114
US 2000-247622P
                     P
                        20001114
US 2000-247630P
                     Р
                        20001114
US 2000-247631P
                     Р
                        20001114
US 2000-247632P
                     Р
                        20001114
                     Р
US 2000-247633P
                        20001114
                     P
                        20001114
US 2000-247556P
                     P
                        20001114
US 2000-247558P
US 2000-247559P
                     Р
                        20001114
US 2000-247560P
                     P
                        20001114
US 2000-247561P
                     Р
                        20001114
US 2000-247594P
                     р
                        20001114
                     Р
                        20001114
US 2000-247595P
                     Р
                        20001114
US 2000-247606P
                     Р
                        20001114
US 2000-247607P
                     р
                        20001114
US 2000-247608P
                     р
                        20001114
US 2000-247609P
                     р
                        20001114
US 2000-247610P
                     Р
                        20001114
US 2000-247611P
                     Р
                        20001114
US 2000-247612P
                     Р
                        20001114
US 2000-247620P
US 2000-247621P
                     Р
                        20001114
US 2000-247634P
                     Р
                        20001114
                     Р
US 2000-247635P
                        20001114
                        20001114
                     Р
US 2000-247698P
                        20001114
                     P
US 2000-247699P
US 2000-247701P
                     Р
                        20001114
                        20001114
US 2000-247702P
                     Р
                        20001114
                     Р
US 2000-247797P
                     Р
US 2000-247798P
                        20001114
                     P
US 2000-247799P
                        20001114
                     P
US 2000-247800P
                        20001114
US 2000-247801P
                     Р
                        20001114
US 2000-247802P
                     Р
                        20001114
                     Р
US 2000-247803P
                        20001114
                     Р
US 2000-247804P
                        20001114
                     W
WO 2001-US26142
                        20010822
```

AB Claimed are compns. comprising a polypeptide and an active agent covalently attached to the polypeptide and a method for delivery of an active agent to a patient by administering the composition to the patient. The peptide is a homopolymer of a naturally occurring amino acid or a heteropolymer of two or more naturally occurring amino acids. In an example, (Glu)n-cephalexin was prepared from Glu(OBut)NCA and cephalexin hydrochloride.

REFERENCE COUNT:

11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 46 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

137:332985

ACCESSION NUMBER:

2002:327387 CAPLUS

DOCUMENT NUMBER: TITLE:

Superior analgesic effect of loxoprofen Na, a prodrug of a non-selective COX inhibitor, over

COX-2 selective inhibitors in rats

AUTHOR (S):

Makino, Mitsuko; Kojima, Takayoshi; Hayakawa, Makiko;

Hiramoto, Kumiko; Mori, Masayoshi

CORPORATE SOURCE:

International Product Management & Medical Affairs

Department, Sankyo Co., Ltd., Tokyo, 103-8426, Japan

SOURCE: Annual Report of Sankyo Research Laboratories (2001),

53, 103-108

CODEN: ASRLEC; ISSN: 1341-741X

PUBLISHER: Sankyo Co., Ltd., Research Institute

DOCUMENT TYPE: Journal LANGUAGE: English

The analgesic effect of loxoprofen Na, a prodrug of a non-selective COX inhibitor, was compared with indomethacin, a non-selective COX inhibitor, and selective COX-2 inhibitors such as celecoxib, rofecoxib and meloxicam by using the Randall-Selitto method in rats. Loxoprofen Na suppressed pain sensation at doses of 0.13 mg/kg and higher, while indomethacin, celecoxib, rofecoxib and meloxicam suppressed this at doses of 8.4, 0.98, 2.4 and 18.3 mg/kg, resp., and higher. Judging from the min. EDs, loxoprofen Na seems to have 7-140 times more potent analgesic action than the selective COX-2 inhibitors and indomethacin. Furthermore, the analgesic activity of loxoprofen Na appeared 15 min after oral administration, which was the shortest latency among the NSAIDs examined Although loxoprofen Na is a prodrug, its rapid oral absorption and conversion to the active form (trans-OH metabolite), which have been already reported, would explain the short latency to the appearance of its analgesic action. This is the first manuscript to report comparison of the analgesic action of a prodrug of a non-selective COX inhibitor with that of selective COX-2 inhibitors in rats. The present data indicates that the analgesic effect of loxoprofen Na is superior to the COX-2 selective inhibitors

REFERENCE COUNT: 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 47 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

examined

2002:276519 CAPLUS

DOCUMENT NUMBER:

136:310188

TITLE:

Treatment of cancer with a prostate specific antigen

(PSA) conjugate and an NSAID compound

INVENTOR(S):

Heimbrook, David C.; Yao, Siu-long

PATENT ASSIGNEE(S):

USA

SOURCE:

U.S. Pat. Appl. Publ., 129 pp.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002042375	A1	20020411	US 2001-896245	20010629
PRIORITY APPLN. INFO.:			US 2000-216217P P	20000705
OTHER SOURCE(S):	MARPAT	136:310188		

The invention relates to methods of treating cancer using a combination of a compound which is a PSA conjugate and a nonsteroidal antiinflammatory agent (NSAID) and to methods of preparing such compns. The PSA conjugate comprises an oligopeptide that is selectively cleaved by PSA and a cytotoxic agent. An example of a PSA conjugate is N-Ac-(4-trans-L-Hyp)-Ala-Ser-Chg-Gln-Ser-Leu-Dox (Dox = doxorubicin, Hyp = hydroxyproline, Chg = cyclohexylglycine) and COX-2 inhibitor 3-phenyl-4-[4-(4-methylsulfonyl)phenyl]-2(5H)furanone is an example of an NSAID compound (syntheses given).

L8 ANSWER 48 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

Page 57

ACCESSION NUMBER:

2001:10616 CAPLUS

DOCUMENT NUMBER:

134:91125

TITLE:

Pharmaceutical compositions containing aldose

reductase inhibitors and selective cyclooxygenase-2

inhibitors

INVENTOR(S): PATENT ASSIGNEE(S): Mylari, Banavara Lakshman Pfizer Products Inc., USA Eur. Pat. Appl., 103 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1064965	A2	20010103	EP 2000-305361	20000626
EP 1064965	A3	20030212		
R: AT, BE, CH,	DE, DK	, ES, FR, GB	, GR, IT, LI, LU, NL,	SE, MC, PT,
IE, SI, LT,	LV, FI	, RO		
US 6555540	B1	20030429	US 2000-602419	20000623
JP 2001031569	A2	20010206	JP 2000-194053	20000628
CA 2313063	AA	20001230	CA 2000-2313063	20000629
BR 2000002957	A	20010130	BR 2000-2957	20000630
PRIORITY APPLN. INFO.:			US 1999-141695P	P 19990630

MARPAT 134:91125 OTHER SOURCE(S):

Pharmaceutical compns. containing aldose reductase inhibitors, a prodrug thereof or a salts and and selective cyclooxygenase-2 inhibitors, a prodrug thereof or salts thereof are disclosed. The compns. are used for the treatment of diabetic complications such as diabetic neuropathy, diabetic nephropathy, diabetic retinopathy and diabetic cardiomyopathy. Hard gelatin capsules contained active ingredients 0.25-100, starch 0.0-650, starch powder 0.0-50, and silicone fluid 350-cSt 0.15 mg/capsules.

ANSWER 49 OF 49 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

1998:202670 CAPLUS

DOCUMENT NUMBER:

128:266249

TITLE:

Diphenyl stilbenes as prodrugs to cyclooxygenase-2

inhibitors, pharmaceutical compositions, and

preparation thereof

INVENTOR(S):

Black, Cameron; Girard, Mario; Guay, Daniel; Wang,

Zhaovin

PATENT ASSIGNEE(S):

Merck Frosst Canada, Inc., Can.

SOURCE:

U.S., 21 pp. CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 5733909	A	19980331	US 1997-784663	19970121
PRIORITY APPLN. INFO.:			US 1997-784663	19970121
OTHER SOURCE(S):	MARPAT	128:266249		

GI

$$R^{1}$$
 R^{2}
 C
 X
 R^{3}

AB Compds. I [X = CH2OH, CHO, CO2R4, CONR42; Y = Me, CH2OR5; R1 = S(0)2Me, S(0)2NH2, etc.; R2, R3 = H, halo, C1-6 alkoxy, etc.; R4 = H, C1-6 alkyl, etc.; R5 = C1-6 alkyl, (substituted) benzyl] are disclosed for the treatment of cyclooxygenase-2 mediated diseases. Also disclosed are pharmaceutical compns. containing I for treatment of cyclooxygenase-2 mediated diseases. Compds. of the invention are useful for treating inflammatory diseases susceptible to treatment with a nonsteroidal antiinflammatory agent. Preparation of selected I is described.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> log h	•	
COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	137.54	332.68
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
	ENTRY	SESSION
CA SUBSCRIBER PRICE	-36.75	-39.75

SESSION WILL BE HELD FOR 60 MINUTES STN INTERNATIONAL SESSION SUSPENDED AT 15:49:01 ON 18 SEP 2006